

N-SUBSTITUTED AZACYCLES

This application claims the benefit of U.S. Provisional Application No. 60/510,568, filed October 10, 2003 and U.S. Provisional Application No. 60/609,533, filed September 13, 2004.

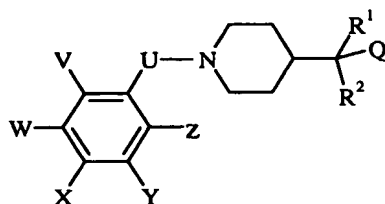
FIELD OF THE INVENTION

The present invention generally relates to novel compounds, processes and intermediates useful in preparing such compounds, compositions containing such compounds and the use of such compounds in controlling insects. In particular, it pertains to N-substituted azacycle derivatives, N-oxides, and agriculturally acceptable salts thereof, compositions of these insecticides, and methods for their use in controlling insects.

BACKGROUND OF THE INVENTION

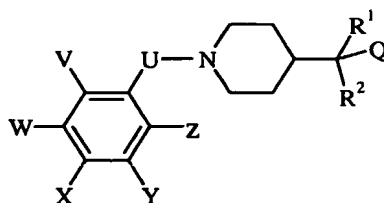
A longstanding worldwide demand exists for new, effective, less costly, and safe means to control pests in agricultural crops, greenhouse crops, nursery crops, ornamentals, turfs, forestry, stored food and fiber products, structures, livestock, households, and public and animal health. Agricultural crop costs incurred by pests exceed billions of dollars annually in decreased crop yields, reduced crop quality and increased harvesting costs. Agricultural crops include wheat, corn, soybeans, potatoes, and cotton to name a few. Soil-borne insects, such as termites and white grubs, cause millions of dollars of damage to structures, turfs and ornamentals. Household pests, such as flies, ants and cockroaches, carry disease and are undesirable in peoples' homes. In addition to these pests, many blood-feeding insects are vectors for pathogenic microorganisms that threaten human and animal health, or are annoying at the least. Insecticides are desired which can control these pests without damaging crops, turfs, ornamentals or structures, and which have no deleterious effects to mammals and other living organisms.

A number of patents disclose a variety of insecticidally active azacycle derivatives. For example, as set forth in United States Patent 5,569,664, compounds of the following structure are reported to be insecticidally active:



where U is selected from $-(CH_2)_n-$ and ethylidene, where n is 1, 2, or 3; Q is selected from hydrogen, hydroxy, sulfhydryl, and fluorine; V is selected from hydrogen, halogen, alkyl, haloalkyl, alkoxy, alkylthio, alkylsulfinyl, alkylsilyloxy, dialkylamino, cyano, nitro, hydroxy, and phenyl; W is selected from hydrogen, halogen, alkyl, haloalkyl, alkoxy, nitro, amino, phenoxy, and phenylalkoxy; X is selected from hydrogen, hydroxy, halogen, alkyl, alkoxyalkyl, alkoxy, cycloalkylalkoxy, haloalkoxy, alkenyloxy, alkynyloxy, alkylsilyloxy, alkylthio, haloalkylthio, cyano, cyanoalkoxy, nitro, amino, monoalkylamino, dialkylamino, alkylaminoalkoxy, alkylcarbonylamino, alkoxycarbonylamino, alkylcarbonyl, alkoxycarbonyl, alkylaminocarbonyl, aminocarbonyloxy, phenyl, phenylalkoxy, phenoxy, and phenoxyalkyl; Y and Z are independently selected from hydrogen and alkoxy; R¹ and R² are independently selected from phenyl substituted with halogen, alkyl, haloalkyl, haloalkoxy, alkoxyalkyl, hydroxy, arylthio, alkoxy, dialkylamino, dialkylaminosulfonyl, hydroxyalkylaminocarbonyl, alkylsulfonyloxy, and haloalkylsulfonyloxy; and the corresponding N-oxides and agriculturally acceptable salts.

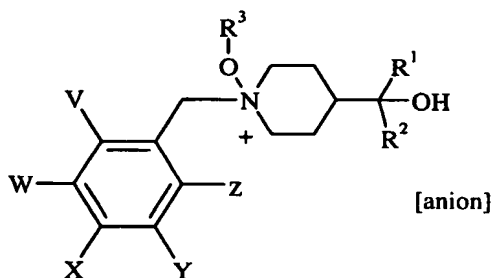
As set forth in United States Patent 5,639,763 compounds of the following structure are reported to be insecticidally active:



where U is selected from $-(CH_2)_n-$ and ethylidene, where n is 1, 2, or 3; Q is selected from hydrogen, hydroxy, sulfhydryl, and fluorine; V is selected from

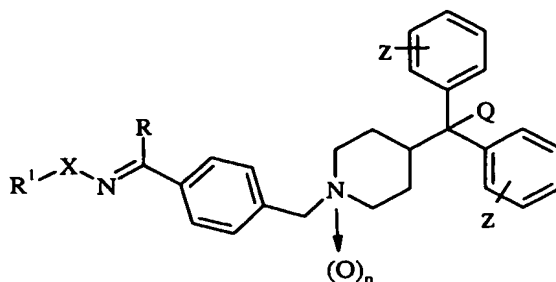
hydrogen, halogen, alkyl, haloalkyl, alkoxy, alkylthio, alkylsulfinyl, alkylsilyloxy, dialkylamino, cyano, nitro, hydroxy, and phenyl; Y and Z are independently selected from hydrogen and alkoxy; W and X taken together is $-\text{OCH}_2\text{CH}_2\text{O}-$, $-\text{CH}_2\text{C}(\text{CH}_3)_2\text{O}-$, $-\text{OC}(\text{CH}_3)_2\text{O}-$, or $-\text{N}=\text{C}(\text{C}_2\text{H}_5)\text{O}-$; R^1 and R^2 are independently selected from phenyl substituted with halogen, alkyl, haloalkyl, haloalkoxy, alkoxyalkyl, hydroxy, arylthio, alkoxy, dialkylamino, dialkylaminosulfonyl, hydroxyalkylaminocarbonyl, alkylsulfonyloxy, and haloalkylsulfonyloxy; and the corresponding N-oxides and agriculturally acceptable salts.

As set forth in United States Patent 5,795,901 compounds of the following structure are reported to be insecticidally active:



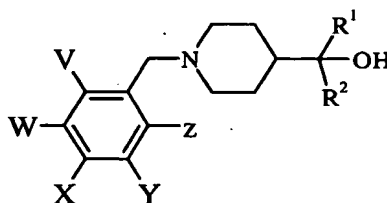
where V, W, Y, and Z are hydrogen; X is alkoxy, cycloalkoxy, alkoxy carbonyl, alkoxy carbonylamino, or a five- or six-membered heteroaryl or heteroaryloxy, each heteroaryl optionally substituted with halogen, cyano, alkyl, haloalkyl, alkoxy, haloalkoxy, alkoxyalkyl, or haloalkoxyalkyl; R^1 and R^2 are independently selected from haloalkyl, phenyl substituted with halogen, haloalkyl, or haloalkoxy; or a five- or six-membered heteroaryl substituted with halogen or alkyl; R^3 is alkyl, haloalkyl, hydroxyalkyl, alkoxyalkyl, dialkylaminoalkyl, alkylaminocarbonyloxyalkyl, alkylthioalkyl, alkylsulfonylalkyl, alkylcarbonyloxyalkyl, alkoxy carbonylalkyl, carboxyalkyl, carboxyarylalkyl, arylcarbonyl, sulfonato, or sulfonatoalkyl, and may bear a negative charge resulting in an inner salt, and a separate anion is chloride, bromide, iodide, or a phenyl, or alkyl sulfate or sulfonate.

As set forth in United States Patent 5,939,438 compounds of the following structure are reported to be insecticidally active:



where R is hydrogen, halogen, alkyl, alkoxy, or dialkylamino; R¹ is
 5 hydrogen, alkyl, haloalkyl, alkoxyalkyl, alkylcarbonyl, , or alkylaminocarbonyl; Q is
 fluoro or hydroxy; X is oxygen or NR²; Z is halogen, haloalkyl, haloalkoxy,
 pentahalothio, haloalkylthio, haloalkylsulfinyl, haloalkylsulfonyl, , or -OCF₂O-
 attached to two adjacent carbon atoms of the phenyl ring; n is 0 or 1; and, when X is
 NR², R² is hydrogen, alkyl, alkylcarbonyl, alkoxyalkyl, or R¹ and R² taken
 10 together may be -C_mH_{2m}-, or -C₂H₄OC₂H₄-, where m is 3-9; and their agriculturally
 acceptable salts.

As set forth in United States Patent 6,017,931 compounds of the following
 structure are reported to be insecticidally active:

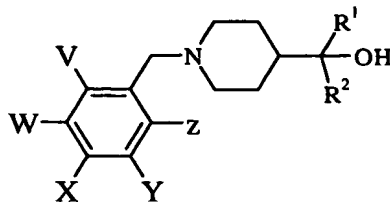


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where V, W, and Z are hydrogen; X is selected from alkoxy, haloalkoxy,
 alkoxyalkyl, cycloalkylalkoxy, halocycloalkylalkoxy, alkoxyalkyl,
 haloalkoxyalkyl, cycloalkylalkoxyalkyl, halocycloalkylalkoxyalkyl,
 20 alkoxyalkoxyalkyl, alkoxyalkoxyalkylamino, haloalkoxyalkoxyalkylamino,
 cycloalkylalkoxyalkoxyalkylamino, halocycloalkylalkoxyalkoxyalkylamino,
 alkylaminocarbonyl, haloalkylaminocarbonyl, cyanoalkoxyalkoxyalkylamino,
 phenylcarbonyl, and phenoxyalkoxyalkyl, each cycloalkyl moiety or phenyl ring
 optionally substituted with halogen; Y is selected from hydrogen or halogen; R¹ and

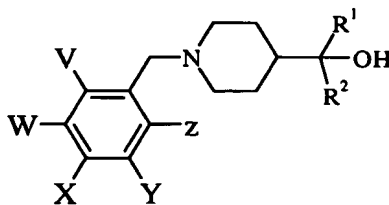
R^2 are independently selected from phenyl or pyridyl, each substituted with haloalkyl, haloalkoxy, or alkylthio, and the corresponding N-oxides and agriculturally acceptable salts.

As set forth in United States Patent 6,030,987 compounds of the following structure are reported to be insecticidally active:



where V, W, Y and Z are hydrogen; X is a five- or six-membered heterocycle optionally substituted with halogen, alkyl, alkoxy, alkoxyalkyl, cyano, aminocarbonyl, haloalkyl, haloalkoxy, or haloalkoxyalkyl; and the heterocycle is optionally connected to the phenyl ring through a -O-, -S-, -(CH₂)_p-, -C(O)-, or -O(CR³R⁴)_q- linkage; R¹ and R² are independently selected from phenyl or pyridyl, each substituted with haloalkyl, or haloalkoxy; R³ and R⁴ are independently selected from hydrogen and methyl; n and p are independently 1, 2, or 3; and q is 1 or 2, and the corresponding N-oxides and agriculturally acceptable salts.

As set forth in United States Patent 6,184,234 compounds of the following structure are reported to be insecticidally active:



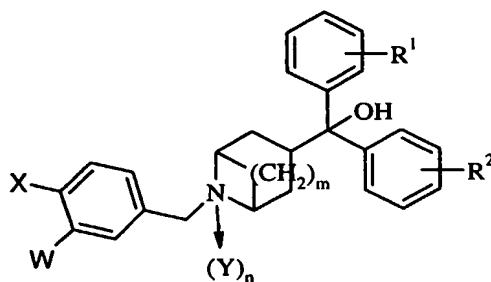
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where V, W, Y and Z are hydrogen; X is a five- or six-membered heterocycle optionally substituted with bromine, chlorine, fluorine, alkyl, alkoxy, alkoxyalkyl, cyano, aminocarbonyl, haloalkyl, haloalkoxy, or haloalkoxyalkyl; and the heterocycle is optionally connected to the phenyl ring through a -O-, -S-, -

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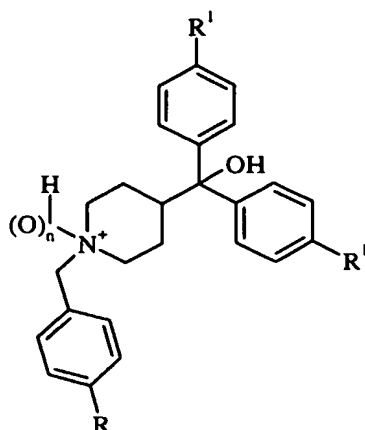
(CH₂)_p-, -C(O)-, or -O(CR³R⁴)_q- linkage; R¹ and R² are independently selected from :
 i) phenyl or pyridyl, each substituted with pentahalothio, haloalkylthio, haloalkylsulfinyl, or haloalkylsulfonyl; ii) phenyl substituted with -OC(M)₂O-,
 5 where M is bromine, chlorine, or fluorine to provide a dihalobenzodioxolyl fused ring; or iii) pyridyl substituted with -OC(M)₂O-, to provide a dihalodioxoleneopyridyl fused ring; R³ and R⁴ are independently selected from hydrogen and methyl; n and p are independently 1, 2, or 3; and q is 1 or 2, and the corresponding N-oxides and agriculturally acceptable salts.

As set forth in United States Statutory Invention Registration H1,838
 10 compounds of the following structure are reported to be insecticidally active:



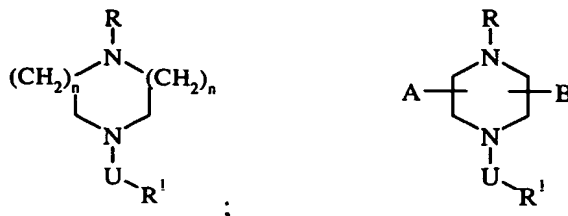
where m is 2 or 3; n is 0 or 1; W is hydrogen or alkoxy; X is hydrogen,
 15 alkoxy, cycloalkylalkoxy, haloalkoxyimino, or a five- or six-membered heteroaryl or heteroaryloxy in which one or more hetero atoms may be optionally substituted with alkyl; R¹ and R² are independently selected from hydrogen, haloalkyl, halothio, or haloalkoxy; and when n is 1, Y represents (a) an N-oxide of the ring nitrogen; or (b) an agriculturally acceptable anionic salt of the ring nitrogen; or (c) forms an OR³
 20 linkage in which R³ is selected from hydrogen, alkyl, alkoxycarbonylalkyl, hydroxycarbonylethyl in association with an agriculturally acceptable anion resulting in an ionic salt, or R³ is an oxycarbonylalkyl group bearing a negative charge resulting in an inner salt.

As set forth in United States Statutory Invention Registration H1,996
 25 photostable, agriculturally acceptable acid salts of an organic or inorganic acid of the following structure are reported to be insecticidally active:



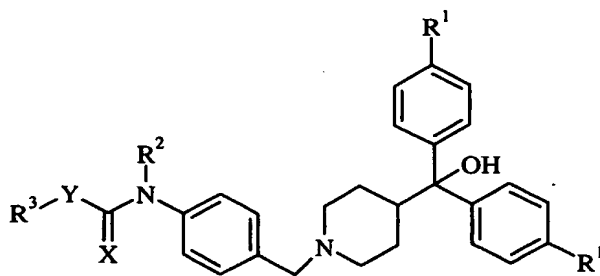
where R is alkoxycarbonyl, alkoxycarbonylamino, cycloalkylalkoxy, 2-alkyl-2H-tetrazol-5-yl, or 2-haloalkyl-2H-tetrazol-5-yl; R¹ is trihaloalkyl, or trihaloalkoxy; n is 0, or 1; and said salt is at least 2.5 times more photostable than its non-ionic parent and is derived from hydrochloric acid, hydrobromic acid, boric acid, phosphoric acid, maleic acid, fumaric acid, phthalic acid, D-glucuronic acid; the sulfonic acid R²SO₃H where R² is alkyl, haloalkyl, hydroxyalkyl, D-10-camphoryl, or phenyl optionally substituted with alkyl or halogen; the carboxylic acid R³CO₂H where R³ is hydrogen, alkyl, trihaloalkyl, carboxyl, phenyl optionally substituted with alkyl or halogen, or pyridyl; the boronic acid R⁴B(OH)₂ where R⁴ is alkyl or phenyl optionally substituted with alkyl or halogen; the phosphonic acid R⁵PO₃H₂ where R⁵ is alkyl, haloalkenyl, or phenyl optionally substituted with alkyl or halogen; the sulfuric acid R⁶OSO₃H where R⁶ is hydrogen or alkyl; or the alkanolic acid X-(CH₂)_qCO₂H where q is 0 to 11, X is halogen, trihaloalkyl, haloalkenyl, cyano, aminocarbonyl, or CO₂R⁷ where R⁷ is hydrogen or alkyl.

As set forth in United States Statutory Invention Registration H2,007 compounds of the following structures are reported to be insecticidally active:



where A and B are independently selected from lower alkyl; U is selected from lower alkylidene, lower alkenylidene, and CH-Z, where Z is selected from hydrogen, lower alkyl, lower cycloalkyl, or phenyl; R is $-\text{CHR}^3\text{R}^4$ where R^3 and R^4 are independently selected from phenyl, optionally substituted with halogen, lower alkyl, lower haloalkyl, lower alkoxy, lower haloalkoxy, lower alkenyl, or phenyl; R^1 is phenyl, naphthyl, tetrazolylphenyl, phenylcyclopropyl, phenoxyphenyl, benzyloxyphenyl, pyridylphenyl, pyridyloxyphenyl, or thiadiazolyloxyphenyl, each optionally substituted with halogen, cyano, hydroxy, lower alkyl, lower haloalkyl, lower alkoxy, amino, lower dialkylamino, nitro, lower haloalkylsulfonyloxy, lower alkylcarbonyloxy, lower alkylcarbonylamino, lower alkoxy carbonyl, lower alkoxyalkoxy carbonyl, lower cycloalkylalkoxy carbonyl, lower alkoxyalkylalkoxy carbonyl, lower alkoxy carbonylamino, alkoxythiocarbonylamino, lower alkyl dithiocarbonylamino, lower dialkyldioxolylalkoxy carbonylamino, or halophenylamino; or lower alkyl substituted with any one of the foregoing cyclic R^1 groups; m is 2 or 3; and n is 1, 2, or 3.

As set forth in unexamined Japanese Patent Application 2002-220372 compounds of the following structures are reported to be insecticidally active:



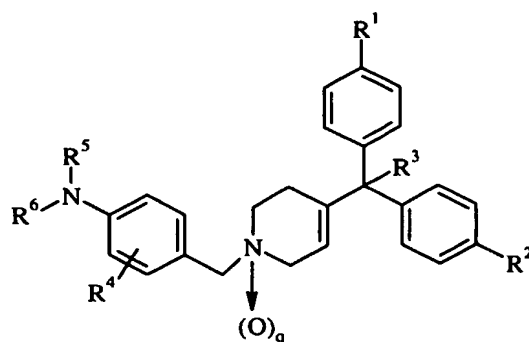
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where R^1 and R^2 are independently selected from hydrogen, halogen, lower alkyl, lower haloalkyl, lower alkoxy, lower haloalkoxy, or lower alkylsulfonyloxy; R^2 is selected from hydrogen, lower alkyl, lower alkenyl, lower alkoxyalkyl, or lower alkylcarbonyl; X and Y are independently oxygen or sulfur; R^3 is selected from lower alkenyl, or lower alkynyl, which are optionally substituted with hydroxy, halogen, lower alkoxy, lower haloalkoxy, lower alkylthio, lower alkylsulfinyl, lower alkylsulfonyl, lower cycloalkyl, lower alkoxyalkoxy, amino, lower alkylamino,

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lower dialkylamino, lower alkoxycarbonyl, nitro, cyano, trimethylsilyl, phenyl, or lower cycloalkenyl; and the corresponding N-oxides and salts.

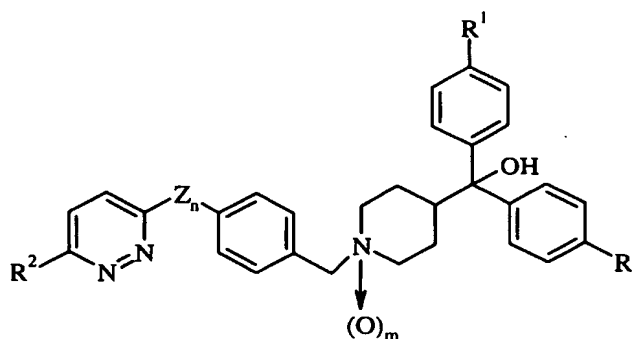
As set forth in PCT Publication WO 02/068392A1 compounds of the following structures are reported to be insecticidally active:



- where R^1 and R^2 are independently selected from halogen, C_1 - C_6 alkyl, halo C_1 - C_6 alkyl, C_1 - C_6 alkoxy, halo C_1 - C_6 alkoxy, $-S(=O)_p-R^9$, or SF_5 ; R^3 is hydrogen, hydroxy, C_1 - C_6 alkoxy, , or $-OC(=O)-C_1$ - C_6 alkyl; R^4 is hydrogen, halogen, C_1 - C_6 alkyl, halo C_1 - C_6 alkyl, C_1 - C_6 alkoxy, halo C_1 - C_6 alkoxy, or $-S(=O)_p-R^9$, or $-SCN$; R^5 and R^6 are independently selected from C_1 - C_{12} alkyl, halo C_1 - C_{12} alkyl, C_2 - C_{12} alkenyl, halo C_2 - C_{12} alkenyl, C_2 - C_{12} alkynyl, halo C_2 - C_{12} alkynyl, C_3 - C_8 cycloalkyl, $-C(=O)-OR^7$, $-C(=S)-OR^8$, $-C(=Y)-ZR^8$, $-S(=O)_p-R^9$, aryl, aryl C_1 - C_6 alkyl, heterocycle, heterocycle C_1 - C_6 alkyl, each substituted in the ring from one to five times independently of one another by halogen, hydroxy, cyano, nitro, C_1 - C_6 alkyl, halo C_1 - C_6 alkyl, C_1 - C_6 alkoxy, halo C_1 - C_6 alkoxy; or in common together with the nitrogen atom to which they are attached to form a heterocyclic ring which is substituted or unsubstituted; Y is oxygen or sulfur; X is a bond, $-NR^{10}$ -, or sulfur; R^7 is C_1 - C_6 alkoxy- C_1 - C_6 alkyl, C_1 - C_6 alkylthio- C_1 - C_6 alkyl, C_1 - C_6 alkylamino- C_1 - C_6 alkyl, C_3 - C_6 alkynyl, C_1 - C_6 alkyl- $S(=O)_p$ - C_1 - C_6 alkyl, C_3 - C_8 cycloalkyl, aryl, aryl- C_1 - C_6 alkyl, heterocyclyl, or heterocyclyl- C_1 - C_6 alkyl each substituted in the ring from one to five times independently of one another by halogen, cyano, nitro, C_1 - C_6 alkyl, halo C_1 - C_6 alkyl, C_1 - C_6 alkoxy, or halo C_1 - C_6 alkoxy; R^8 is C_1 - C_6 alkyl, halo C_1 - C_6 alkyl, C_1 - C_6 alkoxy- C_1 - C_6 alkyl, C_1 - C_6 alkylthio- C_1 - C_6 alkyl, C_2 - C_6 alkenyl, C_3 - C_6 alkynyl, C_1 - C_6 alkyl- $S(=O)_p$ - C_1 - C_6 alkyl, C_3 - C_8 cycloalkyl, aryl, aryl- C_1 - C_6 alkyl, heterocyclyl,

or heterocyclyl-C₁-C₆alkyl, or is C₃-C₈cycloalkyl, aryl, aryl-C₁-C₆alkyl, heterocyclyl, or heterocyclyl-C₁-C₆alkyl each substituted in the ring from one to five times independently of one another by halogen, cyano, nitro, C₁-C₆alkyl, haloC₁-C₆alkyl, C₁-C₆alkoxy, or haloC₁-C₆alkoxy; R⁹ is C₁-C₆alkyl, C₃-C₈cycloalkyl, haloC₁-C₆alkyl, or benzyl; R¹⁰ is hydrogen, C₁-C₆alkyl, C₃-C₈cycloalkyl, haloC₁-C₆alkyl, or benzyl; p is 0, 1, or 2; q is 0 or 1; and, where appropriate, E/Z isomers, E/Z isomer mixtures and/or tautomers, each in free form or in salt form.

As set forth in PCT Publication WO 200020409A1 compounds of the following structures are reported to be insecticidally active:

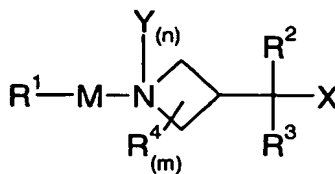


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where R¹ is halo, C₁-C₄alkyl, C₁-C₄alkoxy, C₁-C₄haloalkyl, C₁-C₄haloalkoxy; R² is hydrogen, hydroxyl, halo, C₁-C₄alkyl, C₁-C₄alkoxy, C₁-C₄alkoxycarbonyl, C₁-C₄alkylthio, C₁-C₄alkylsulfonyl, optionally substituted phenyl or carbamoyl; Z is O or S(O)_p, p is 0 or 2; and m and n are 0 or 1.

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As set forth in PCT Publication WO 03/022808A1 compounds of the following structures are reported to be insecticidally active:



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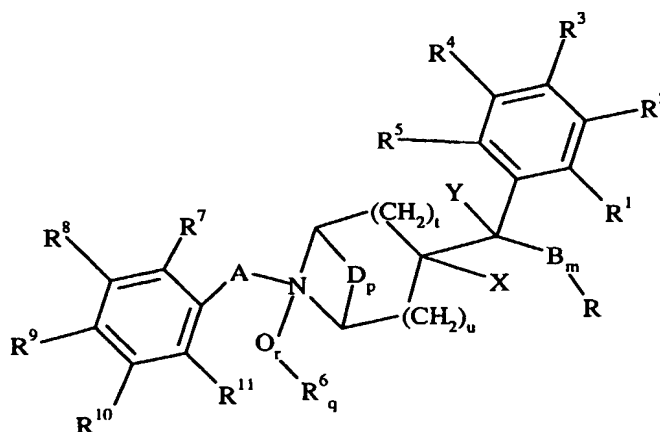
where R¹ is an aryl or heteroaryl that is optionally identically or differently substituted once or several times; R² and R³ are independently selected from aryl or heteroaryl that is optionally identically or differently substituted once or several times, whereby both groups can also be bridged by a common substituent; M is

optionally substituted $(\text{CH}_2)_l$, where l is 1, 2 or 3, CO or $-\text{HNC}(\text{O})$; X is H, OH, halogen, OR^4 or CN; Y is (O), H, OH, OR^4 , R^4 ; (in the last four groups, in which nitrogen has a positive charge, in combination with a corresponding anion); R^4 is identical or different and represents C_1 - C_4 alkyl, C_1 - C_4 alkanoyl, C_1 - C_4 haloalkyl; m is 0, 1, 2, 3 or 4; and n is 0 or 1.

There is no disclosure or suggestion in any of the citations set forth above of the azacycle derivatives of the present invention.

SUMMARY OF THE INVENTION

In accordance with the present invention, it has now been found that N-substituted azacycle derivatives of Formula I and salts thereof having a substituent X as indicated in the Formula have improved insecticidal activity. The compounds of formula I are represented by the following general formula I:



I

wherein;

m , q , r , t and u are independently selected from 0 or 1; and p is 0, 1, 2, or 3;

X is selected from halogen, hydroxyl, hydroxyalkyl, alkyl, alkoxy, haloalkyl, haloalkoxy, thio, alkylthio, acetoxyalkyl, azidoalkyl, aminoalkyl, acetylaminalkyl, alkylsulfonyl, alkylsulfoxy, pentahalothio, cyano, nitro, acetyloxy, alkylsulfonyloxy, alkylcarbonyl, alkoxycarbonyl, aryl, or aryloxy;

Y is selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyl, hydroxyalkyl, alkoxy, haloalkoxy, thio, alkylthio, pentahalothio, cyano, nitro, alkylsulfonyl, alkylsulfoxy, alkylsulfonyloxy, alkylcarbonyl, alkoxycarbonyl, aryl, or aryloxy;

5 or

X and Y taken together with $-\text{OCR}^{12}\text{R}^{13}\text{O}-$, form a 1,3-dioxolane ring;

where

R^{12} and R^{13} are independently selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyalkyl, alkoxy, haloalkoxy, alkylthio, cyano, nitro, alkylsulfonyloxy, alkylcarbonyl, alkoxycarbonyl, aryl, or aryloxy;

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or

R^{12} and R^{13} taken together with $(=\text{O})$, form 1,3-dioxol-2-one ring;

R^1 , R^2 , R^3 , R^4 , and R^5 are independently selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyl, hydroxyalkyl, alkoxy, haloalkoxy, thio, alkylthio, haloalkylthio, pentahalothio, cyano, nitro, alkylsulfonyl, haloalkylsulfonyl, alkylsulfinyl, haloalkylsulfinyl, alkylsulfonyloxy, haloalkylsulfonyloxy, alkylcarbonyl, alkoxycarbonyl, dialkoxyalkylcarbonyl, alkoxycarbonylamino, alkylaminoxyalkyl, alkoxyiminoalkyl, alkenyloxyiminoalkyl, aryl, aryloxy, dioxanyl, dioxolanyl or either of R^1 and R^2 , or R^2 and R^3 , or R^3 and R^4 , or R^4 and R^5 taken together with $-\text{OC}(\text{R}^{19})_2\text{O}-$, $-\text{OC}(\text{R}^{19})_2(\text{R}^{19})_2\text{O}-$, $-\text{OC}(\text{R}^{19})_2(\text{R}^{19})_2-$, $-\text{OC}(\text{R}^{19})=\text{N}-$, or $-\text{SC}(\text{R}^{19})=\text{N}-$, forming a benzo-fused ring, where R^{19} is hydrogen, halogen, alkyl or haloalkyl; and, wherein at least one of R^1 , R^2 , R^3 , R^4 , and R^5 is other than hydrogen;

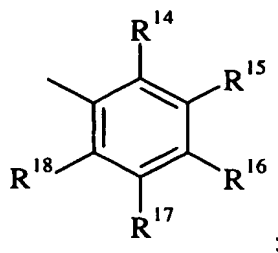
R^7 , R^8 , R^9 , R^{10} , and R^{11} are independently selected from hydrogen, halogen, alkyl, hydroxy, hydroxyalkyl, hydroxyalkoxy, alkoxy, alkoxyalkyl, alkoxyiminoalkyl, haloalkoxyiminoalkyl, cyanoalkoxyiminoalkyl, cyanoiminothioalkylamino, alkenyloxyiminoalkyl, alkynyloxyiminoalkyl, cycloalkoxy, cycloalkylalkoxy, phenoxy, alkoxycarbonylphenoxy, alkoxyalkoxy, alkoxyalkoxyalkoxy, alkylthio, alkylsulfonyl, alkylaminosulfonyl, dialkylaminosulfonyl, cycloalkylaminosulfonyl, alkenyloxy, alkynyloxy, haloalkenyloxy, alkylsulfonyloxy, optionally substituted arylalkoxy, cyano, nitro, amino, alkylamino, alkylcarbonylamino,

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- alkoxycarbonylamino, cycloalkylalkoxycarbonylamino, alkenyloxycarbonylamino, alkynyloxycarbonylamino, haloalkylcarbonylamino, alkoxyalkoxycarbonylamino, alkoxycarbonylamino, alkoxycarbonyloxy, alkenylaminocarbonyloxy, alkynylaminocarbonyloxy, (alkyl)(alkoxycarbonyl)amino, alkylsulfonylamino, optionally substituted (heteroaryl)(alkoxycarbonyl)amino, optionally substituted arylcarbonylamino, alkoxycarbonyl, alkylaminocarbonyloxy, alkylaminocarbonylamino, dialkylaminocarbonylamino, alkylamino(thiocarbonyl)amino, dialkylphosphoroureydyl, acetoxyalkoxy, sulfonyloxyalkoxy, dialkoxyalkoxy, trialkoxyalkoxy, dialkoxyalkylacetal, trialkoxymethylorthoester, cyclic acetal, optionally substituted cyclic acetal, optionally substituted thienyl, optionally substituted 1,3-thiazolylalkoxy, optionally substituted aryl, optionally substituted aryloxy, optionally substituted aryloxyalkyl, optionally substituted arylaminocarbonyloxy, optionally substituted arylalkoxycarbonylamino, optionally substituted heteroaryl, optionally substituted heteroaryloxy, optionally substituted pyrrolyl, optionally substituted pyrazolyl, optionally substituted pyrazinyloxy, optionally substituted cycloalkylcarbonylamino, optionally substituted 1,3-oxazoliny, optionally substituted 1,3-oxazolinyloxy, optionally substituted 1,3-oxazolinylamino, optionally substituted 1,2,4-triazolyl, optionally substituted 1,2,3-thiadiazolyl, optionally substituted 1,2,5-thiadiazolyl, optionally substituted 1,2,5-thiadiazolyloxy, optionally substituted 2H-tetrazolyl, optionally substituted pyridyl, optionally substituted pyridyloxy, optionally substituted pyridylamino, optionally substituted pyrimidinyl, optionally substituted pyrimidinyloxy, optionally substituted 3,4,5,6-tetrahydropyrimidinyloxy, optionally substituted pyridazinyloxy, or optionally substituted 1,2,3,4-tetrahydronaphthalenyl, wherein the optional substituent is selected from one or more of halogen, alkyl, haloalkyl, alkoxy, cyano, nitro, amino, alkylcarbonyl, alkoxycarbonyl, alkoxyiminoalkyl, dialkylacetal, alkylthiol, alkylsulfoxide, or alkoxycarbonylamino; and, wherein at least one of R⁷, R⁸, R⁹, R¹⁰, and R¹¹ is other than hydrogen;
- R is alkyl, cycloalkyl, alkenyl, alkoxycarbonyl, optionally substituted pyrid-2-yl wherein the optional substituent is selected from hydrogen, halogen, haloalkoxy or haloalkyl,

or

substituted phenyl have the following structure,



- 5 where
- R^{14} , R^{15} , R^{16} , R^{17} , and R^{18} are independently selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyl, hydroxyalkyl, alkoxy, haloalkoxy, thio, alkylthio, haloalkylthio, pentahalothio, cyano, nitro, alkylsulfonyl, haloalkylsulfonyl, alkylsulfinyl, haloalkylsulfinyl, alkylsulfonyloxy, haloalkylsulfonyloxy, alkylcarbonyl, alkoxy carbonyl, dialkoxyalkylcarbonyl, alkoxy carbonylamino, alkylaminoxyalkyl, alkoxyiminoalkyl, alkenyloxyiminoalkyl, aryl, aryloxy, dioxanyl, dioxolanyl or either of R^{14} and R^{15} , or R^{15} and R^{16} , or R^{16} and R^{17} , or R^{17} and R^{18} taken together with $-\text{OC}(\text{R}^{19})_2\text{O}-$, $-\text{OC}(\text{R}^{19})_2(\text{R}^{19})_2\text{O}-$, $-\text{OC}(\text{R}^{19})_2(\text{R}^{19})_2-$, $-\text{OC}(\text{R}^{19})=\text{N}-$, or $-\text{SC}(\text{R}^{19})=\text{N}-$, forming a benzo-fused ring, where R^{19} is hydrogen, halogen, alkyl or haloalkyl; and, wherein at least one of R^{14} , R^{15} , R^{16} , R^{17} , and R^{18} is other than hydrogen;

- A is selected from $-\text{CH}_2-$, $-\text{CH}_2\text{CH}_2-$, $-\text{CH}_2\text{CH}_2\text{CH}_2-$, $-\text{OCH}_2\text{CH}_2-$, $-\text{OCH}_2\text{CH}_2\text{CH}_2-$, $-\text{OCH}_2\text{CH}_2\text{CH}_2\text{CH}_2-$, $-\text{OCH}_2\text{CH}(\text{OH})\text{CH}_2-$, $-\text{NHCH}_2\text{CH}_2-$, $-\text{N}(\text{CH}_3)\text{CH}_2\text{CH}_2-$, $-\text{N}[\text{C}(=\text{O})\text{CH}_3]\text{CH}_2\text{CH}_2-$, or $-\text{N}[\text{C}(=\text{O})\text{OCH}_3]\text{CH}_2\text{CH}_2-$;

B is selected from $-\text{O}-$, $-\text{S}-$, $-\text{CH}_2\text{O}-$, $-\text{OCH}_2-$, $-\text{OC}(=\text{O})\text{NH}-$, $-\text{OC}(=\text{O})\text{O}-$, or $-\text{NHSO}_2-$;

- 25 when p is 1, 2, or 3;

D-is $-\text{CH}_2-$;

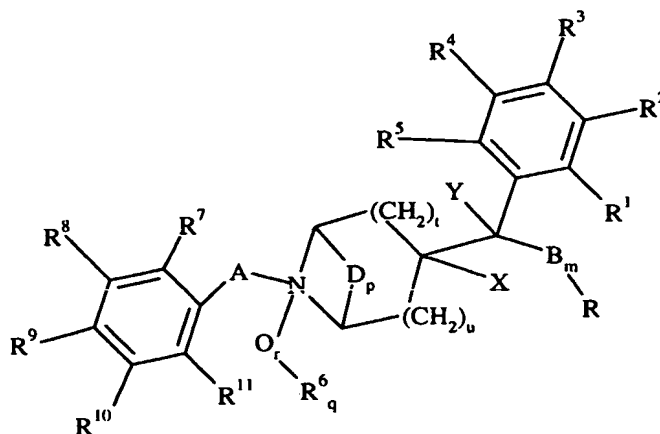
R^6 is selected from alkyl, haloalkyl, hydroxyalkyl, alkoxyalkyl, dialkylaminoalkyl, alkylaminocarbonyloxyalkyl, alkylthioalkyl, alkylsulfonylalkyl, alkylcarbonyloxyalkyl, alkoxycarbonylalkyl, carboxyalkyl, arylalkyl, arylcarbonyl, sulfonato, or sulfonatoalkyl, and may bear a negative charge resulting in an inner salt; and a separate ion is chloride, bromide, iodide, or an alkyl or phenyl sulfate or sulfonate;
 5 and
 agriculturally-acceptable salts thereof.

The present invention is also directed to compositions containing an insecticidally effective amount of at least one of a compound of formula I, and optionally, an effective amount of at least one of a second compound, with at least one agriculturally acceptable extender or adjuvant.

The present invention is also directed to methods of controlling insects, where control is desired, which comprise applying an insecticidally effective amount of the above composition to the locus of crops, or other areas where insects are present or are expected to be present. Other aspects of the present invention will become apparent.

DETAILED DESCRIPTION OF THE INVENTION

20 One aspect of the present invention relates to certain new and useful compounds, namely certain novel N-substituted azacycle derivatives as depicted in general formula I:



I

wherein;

m, q, r, t and u are independently selected from 0 or 1; and p is 0, 1, 2, or 3;

5 X is selected from halogen, hydroxyl, hydroxyalkyl, alkyl, alkoxy, haloalkyl, haloalkoxy, thio, alkylthio, acetoxyalkyl, azidoalkyl, aminoalkyl, acetylaminoalkyl, alkylsulfonyl, alkylsulfoxy, pentahalothio, cyano, nitro, acetyloxy, alkylsulfonyloxy, alkylcarbonyl, alkoxycarbonyl, aryl, or aryloxy;

10 Y is selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyl, hydroxyalkyl, alkoxy, haloalkoxy, thio, alkylthio, pentahalothio, cyano, nitro, alkylsulfonyl, alkylsulfoxy, alkylsulfonyloxy, alkylcarbonyl, alkoxycarbonyl, aryl, or aryloxy;

or

X and Y taken together with $-\text{OCR}^{12}\text{R}^{13}\text{O}-$, form a 1,3-dioxolane ring;

15 where

R^{12} and R^{13} are independently selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyalkyl, alkoxy, haloalkoxy, alkylthio, cyano, nitro, alkylsulfonyloxy, alkylcarbonyl, alkoxycarbonyl, aryl, or aryloxy;

or

20 R^{12} and R^{13} taken together with $(=\text{O})$, form 1,3-dioxol-2-one ring;

R^1 , R^2 , R^3 , R^4 , and R^5 are independently selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyl, hydroxyalkyl, alkoxy, haloalkoxy, thio, alkylthio, haloalkylthio, pentahalothio, cyano, nitro, alkylsulfonyl, haloalkylsulfonyl, alkylsulfinyl, haloalkylsulfinyl, alkylsulfonyloxy, haloalkylsulfonyloxy, alkylcarbonyl, alkoxycarbonyl, dialkoxyalkylcarbonyl, alkoxycarbonylamino, alkylaminoxalkyl, alkoxyiminoalkyl, alkenyloxyiminoalkyl, aryl, aryloxy, dioxanyl, dioxolanyl or either of R^1 and R^2 , or R^2 and R^3 , or R^3 and R^4 , or R^4 and R^5 taken together with $-\text{OC}(\text{R}^{19})_2\text{O}-$, $-\text{OC}(\text{R}^{19})_2(\text{R}^{19})_2\text{O}-$, $-\text{OC}(\text{R}^{19})_2(\text{R}^{19})_2-$, $-\text{OC}(\text{R}^{19})=\text{N}-$, or $-\text{SC}(\text{R}^{19})=\text{N}-$, forming a benzo-fused ring, where R^{19} is hydrogen, halogen, alkyl or haloalkyl; and,

30 wherein at least one of R^1 , R^2 , R^3 , R^4 , and R^5 is other than hydrogen;

R⁷, R⁸, R⁹, R¹⁰, and R¹¹ are independently selected from hydrogen, halogen, alkyl, hydroxy, hydroxyalkyl, hydroxyalkoxy, alkoxy, alkoxyalkyl, alkoxyiminoalkyl, haloalkoxyiminoalkyl, cyanoalkoxyiminoalkyl, cyanoiminothioalkylamino, alkenyloxyiminoalkyl, alkynyloxyiminoalkyl, cycloalkoxy, cycloalkylalkoxy, phenoxy, alkoxyacetylphenoxy, alkoxyalkoxy, alkoxyalkoxyalkoxy, alkylthio, alkylsulfonyl, alkylaminosulfonyl, dialkylaminosulfonyl, cycloalkylaminosulfonyl, alkenyloxy, alkynyloxy, haloalkenyloxy, alkylsulfonyloxy, optionally substituted arylalkoxy, cyano, nitro, amino, alkylamino, alkylcarbonylamino, alkoxyacetylamino, cycloalkylalkoxyacetylamino, alkenyloxyacetylamino, alkynyloxyacetylamino, haloalkylcarbonylamino, alkoxyalkoxyacetylamino, alkoxyacetylamino, alkoxyacetyloxy, alkenylaminocarbonyloxy, alkynylaminocarbonyloxy, (alkyl)(alkoxyacetyl)amino, alkylsulfonylamino, optionally substituted (heteroaryl)(alkoxyacetyl)amino, optionally substituted arylcarbonylamino, alkoxyacetyl, alkylaminocarbonyloxy, alkylaminocarbonylamino, dialkylaminocarbonylamino, alkylamino(thiocarbonyl)amino, dialkylphosphoroureydyl, acetoxyalkoxy, sulfonyloxyalkoxy, dialkoxyalkoxy, trialkoxyalkoxy, dialkoxyalkylacetal, trialkoxymethylorthoester, cyclic acetal, optionally substituted cyclic acetal, optionally substituted thienyl, optionally substituted 1,3-thiazolylalkoxy, optionally substituted aryl, optionally substituted aryloxy, optionally substituted aryloxyalkyl, optionally substituted arylaminocarbonyloxy, optionally substituted arylalkoxyacetylamino, optionally substituted heteroaryl, optionally substituted heteroaryloxy, optionally substituted pyrrolyl, optionally substituted pyrazolyl, optionally substituted pyrazinyloxy, optionally substituted cycloalkylcarbonylamino, optionally substituted 1,3-oxazolinyll, optionally substituted 1,3-oxazolinyloxy, optionally substituted 1,3-oxazolinyllamino, optionally substituted 1,2,4-triazolyl, optionally substituted 1,2,3-thiadiazolyl, optionally substituted 1,2,5-thiadiazolyl, optionally substituted 1,2,5-thiadiazolyloxy, optionally substituted 2H-tetrazolyl, optionally substituted pyridyl, optionally substituted pyridyloxy, optionally substituted pyridylamino, optionally substituted pyrimidinyl, optionally substituted pyrimidinylloxy, optionally substituted 3,4,5,6-tetrahydropyrimidinylloxy, optionally substituted pyridazinylloxy, or optionally substituted 1,2,3,4-tetrahydronaphthalenyl,

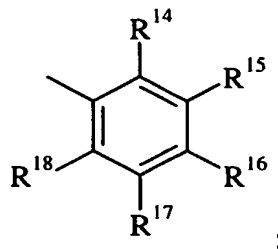
wherein the optional substituent is selected from one or more of halogen, alkyl, haloalkyl, alkoxy, cyano, nitro, amino, alkylcarbonyl, alkoxy carbonyl, alkoxyiminoalkyl, dialkylacetal, alkylthiol, alkylsulfoxide, or alkoxy carbonylamino; and, wherein at least one of R^7 , R^8 , R^9 , R^{10} , and R^{11} is other than hydrogen;

5

R is alkyl, cycloalkyl, alkenyl, alkoxy carbonyl, optionally substituted pyrid-2-yl wherein the optional substituent is selected from hydrogen, halogen, haloalkoxy or haloalkyl,

or

10 substituted phenyl have the following structure,



where

R^{14} , R^{15} , R^{16} , R^{17} , and R^{18} are independently selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyl, hydroxyalkyl, alkoxy, haloalkoxy, thio, alkylthio, haloalkylthio, pentahalothio, cyano, nitro, alkylsulfonyl, haloalkylsulfonyl, alkylsulfinyl, haloalkylsulfinyl, alkylsulfonyloxy, haloalkylsulfonyloxy, alkylcarbonyl, alkoxy carbonyl, dialkoxyalkylcarbonyl, alkoxy carbonylamino, alkylaminoxyalkyl, alkoxyiminoalkyl, alkenyloxyiminoalkyl, aryl, aryloxy, dioxanyl, dioxolanyl or either of R^{14} and R^{15} , or R^{15} and R^{16} , or R^{16} and R^{17} , or R^{17} and R^{18} taken together with $-OC(R^{19})_2O-$, $-OC(R^{19})_2(R^{19})_2O-$, $-OC(R^{19})_2(R^{19})_2-$, $-OC(R^{19})=N-$, or $-SC(R^{19})=N-$, forming a benzo-fused ring, where R^{19} is hydrogen, halogen, alkyl or haloalkyl; and, wherein at least one of R^{14} , R^{15} , R^{16} , R^{17} , and R^{18} is other than hydrogen;

25

A is selected from $-CH_2-$, $-CH_2CH_2-$, $-CH_2CH_2CH_2-$, $-OCH_2CH_2-$, $-OCH_2CH_2CH_2-$, $-OCH_2CH_2CH_2CH_2-$, $-OCH_2CH(OH)CH_2-$, $-NHCH_2CH_2-$, $-N(CH_3)CH_2CH_2-$, $-N[C(=O)CH_3]CH_2CH_2-$, or $-N[C(=O)OCH_3]CH_2CH_2-$;

B is selected from -O-, -S-, -CH₂O-, -OCH₂-, -OC(=O)NH-, -OC(=O)O-, or -NHSO₂-;

5 when p is 1, 2, or 3;

D is -CH₂-;

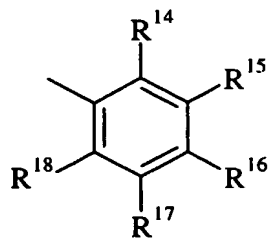
R⁶ is selected from alkyl, haloalkyl, hydroxyalkyl, alkoxyalkyl, dialkylaminoalkyl, alkylaminocarbonyloxyalkyl, alkylthioalkyl, alkylsulfonylalkyl, alkylcarbonyloxyalkyl, alkoxycarbonylalkyl, carboxyalkyl, arylalkyl, arylcarbonyl, sulfonato, or sulfonatoalkyl, and may bear a negative charge resulting in an inner salt; and a separate ion is chloride, bromide, iodide, or an alkyl or phenyl sulfate or sulfonate;

and

15 agriculturally-acceptable salts thereof.

Within the scope set forth above, preferred compounds of the present invention are those of formula I wherein m, q and p are 0; t and u are 1; A is -CH₂-; X is selected from halogen, hydroxyl or alkoxy; Y is selected from hydrogen, halogen or hydroxyl; R¹, R², R³ and R⁴ are independently selected from hydrogen, halogen, alkyl, alkoxy, haloalkyl, haloalkoxy, -CH₂(OH)CH₃, -CH=NOC₂H₅, 1,3-dioxolan-2-yl, or R² and R³ taken together with -OCF₂O-; R⁵ is hydrogen; R⁷, R¹⁰ and R¹¹ are hydrogen; R⁸ is selected from hydrogen, halogen, alkyl or alkoxy; R⁹ is selected from alkoxy, alkoxyalkoxy, alkoxyalkoxyalkoxy, cyclopropylmethoxy, 2-halophenoxy, 3-halophenoxy, 4-halophenoxy, pyrimidin-2-yl, pyrid-2-yl, 3-halo-pyrid-2-yl, 3-alkyl-pyrid-2-yloxy, 4-alkyl-pyrid-2-yloxy, 5-alkyl-pyrid-2-yloxy, 6-alkyl-pyrid-2-yloxy, 3-halo-pyrid-2-yloxy, 3-trihaloalkyl-pyrid-2-yloxy, 3-cyano-pyrid-2-yloxy, 5-cyano-pyrid-2-yloxy, 6-dialkoxyalkyl-pyrid-2-yloxy, pyrid-2-yloxy, CO₂CH(CH₃)₂, -CH=NOCH₃, -CH=NOC₂H₅, -CH=NOCH₂CF₃, -CH=NOCH₂CH=CH₂, -CH=NOCH₂CN, -CH=NOCH(CH₃)₂, -CH=NOCH₂C≡CH, -CH=NOCH₂CH₂F, -CH=NOCH₂CH₂OCH₃, -CH=NOCH₂OC₂H₅, -CH=NOCH₂CH₂OCH₂CH₂OCH₃, -NHCO₂CH₃, -NHCO₂C₂H₅, -NHCO₂CH(CH₃)₂, -NHCO₂CH₂-c-C₃H₅, -CH(OH)C₆H₅-p-Cl, -

OC(=O)NHCH₃, -OC(=O)NHC₂H₅, -OC(=O)NHCH(CH₃)₂, -NHC(SCH₃)=NCN, pyrimidin-2-yloxy, 6-halo-pyridazin-3yloxy, 6-alkoxy-pyridazin-3yloxy, 6-alkyl-pyridazin-3yloxy, 2-alkyl-2H-tetrazol-5-yl, 1,3-dioxan-2-yl or 5,5-dialkyl-1,3-dioxan-2-yl; and R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, and R¹⁸,



5

where

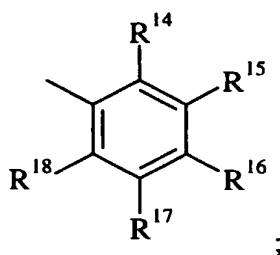
R¹⁴, R¹⁵, R¹⁶ and R¹⁷ are independently selected from halogen, haloalkyl, haloalkoxy or R¹⁵ and R¹⁶ taken together with -OCF₂O-; and R¹⁸ is hydrogen.

Within the scope set forth above, more preferred compounds of the present invention are those of formula I wherein X is selected from halogen, -CO₂C₂H₅ or hydroxyl; and R⁹ is selected from -OC₂H₅, -OC₃H₇, -OCH(CH₃)₂, -OCH₂CH₂OCH₃, -OCH₂CH₂CH₂OCH₃, cyclopropylmethoxy, 2-chlorophenoxy, 3-chlorophenoxy, 4-chlorophenoxy, pyrimidin-2-yl, pyrid-2-yl, 3-chloro-pyrid-2-yl, 3-methyl-pyrid-2-yloxy, 4-methyl-pyrid-2-yloxy, 5-methyl-pyrid-2-yloxy, 6-methyl-pyrid-2-yloxy, 3-chloro-pyrid-2-yloxy, 3-trifluoromethyl-pyrid-2-yloxy, 3-cyano-pyrid-2-yloxy, 5-cyano-pyrid-2-yloxy, 6-dimethoxymethyl-pyrid-2-yloxy, pyrid-2-yloxy, -CO₂CH(CH₃)₂, -CH=NOCH₃, -CH=NOC₂H₅, -CH=NOCH₂CF₃, -CH=NOCH₂CH=CH₂, -CH=NOCH₂CN, -CH=NOCH(CH₃)₂, -CH=NOCH₂C≡CH, -CH=NOCH₂CH₂F, -CH=NOCH₂CH₂OCH₃, -CH=NOCH₂OC₂H₅, -CH=NOCH₂CH₂OCH₂CH₂OCH₃, -NHCO₂CH₃, -NHCO₂C₂H₅, -NHCO₂CH(CH₃)₂, -NHCO₂CH₂-*c*-C₃H₅, -CH(OH)C₆H₅-*p*-Cl, -OC(=O)NHCH₃, -OC(=O)NHC₂H₅, -OC(=O)NHCH(CH₃)₂, -NHC(SCH₃)=NCN, pyrimidin-2-yloxy, 6-chloro-pyridazin-3yloxy, 6-methoxy-pyridazin-3yloxy, 6-methyl-pyridazin-3yloxy, 2-methyl-2H-tetrazol-5-yl, 2-ethyl-2H-tetrazol-5-yl, 1,3-dioxan-2-yl or 5,5-dimethyl-1,3-dioxan-2-yl.

25

Within the scope set forth above, even more preferred compounds of the present invention are those of formula I wherein X is selected from fluorine, -CO₂C₂H₅ or hydroxyl; Y is selected from hydrogen, fluorine, chlorine or hydroxyl;

R^1 , R^2 , R^3 and R^4 are independently selected from hydrogen, halogen, alkyl, tert-butyl, methoxy, trifluoromethyl, difluoromethoxy, trifluoromethoxy, -OCF₂CHFCF₃, -CH₂(OH)CH₃, -CH=NOC₂H₅, 1,3-dioxolan-2-yl, or R^2 and R^3 taken together with -OCF₂O-; R^8 is hydrogen; R^9 is selected from -OCH₂CH₂OCH₃,
 5 -CH=NOCH₃, -CH=NOC₂H₅, -CH=NOCH₂CN, -CH=NOCH₂CH₂OCH₃, -NHCO₂CH(CH₃)₂, -OC(=O)NHCH(CH₃)₂, pyrimidin-2-yl, pyrid-2-yl, 3-chloro-pyrid-2-yl, 3-methyl-pyrid-2-yloxy, 4-methyl-pyrid-2-yloxy, 5-methyl-pyrid-2-yloxy, 6-methyl-pyrid-2-yloxy, 3-chloro-pyrid-2-yloxy, 3-trifluoromethyl-pyrid-2-yloxy, 3-cyano-pyrid-2-yloxy, 5-cyano-pyrid-2-yloxy, 6-dimethoxymethyl-pyrid-2-yloxy,
 10 yloxy, pyrid-2-yloxy, pyrimidin-2-yloxy, 6-chloro-pyridazin-3-yloxy, 6-methoxy-pyridazin-3-yloxy or 6-methyl-pyridazin-3-yloxy; and R is phenyl substituted with R^{14} , R^{15} , R^{16} , R^{17} , and R^{18} ,

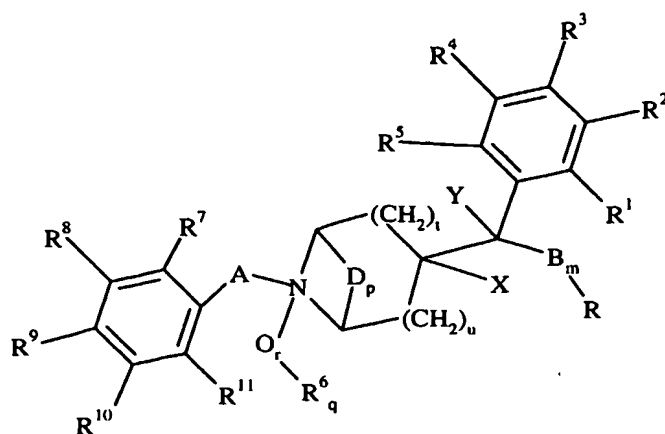


where

15 R^{14} , R^{15} , R^{16} and R^{17} are independently selected from fluorine, chlorine, trifluoromethyl, difluoromethoxy, trifluoromethoxy, -OCF₂CHFCF₃ or R^{15} and R^{16} taken together with -OCF₂O-.

Within the scope set forth above, most preferred compounds of the present invention are those of formula I wherein X is hydroxyl; Y is hydrogen; R^3 is haloalkoxy; R^9 is selected -OCH₂CH₂OCH₃, -CH=NOCH₃, -CH=NOC₂H₅, -CH=NOCH₂CN, -CH=NOCH₂CH₂OCH₃, -NHCO₂CH(CH₃)₂, -OC(=O)NHCH(CH₃)₂, pyrid-2-yloxy, pyrid-2-yl, 3-cyano-pyrid-2-yloxy, 5-methyl-pyrid-2-yloxy, pyrimidin-2-yloxy, pyrimidin-2-yl, 6-chloro-pyridazin-3-yloxy or 6-methoxy-pyridazin-3-yloxy; and R^{16} is haloalkoxy.

25 An embodiment of the present invention is a compound of formula I:

**I**

wherein;

- 5 m, q and r are independently selected from 0 or 1; t and u are 1; and p is 0;

X is selected from halogen, hydroxyl, hydroxyalkyl, alkyl, alkoxy, haloalkyl, haloalkoxy, thio, alkylthio, acetoxyalkyl, azidoalkyl, aminoalkyl, acetylaminalkyl, alkylsulfonyl, alkylsulfoxy, pentahalothio, cyano, nitro, acetyloxy, alkylsulfonyloxy, 10 alkylcarbonyl, alkoxycarbonyl, aryl, or aryloxy;

Y is selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyl, hydroxyalkyl, alkoxy, haloalkoxy, thio, alkylthio, pentahalothio, cyano, nitro, alkylsulfonyl, alkylsulfoxy, alkylsulfonyloxy, alkylcarbonyl, alkoxycarbonyl, aryl, or aryloxy;

15 or

X and Y taken together with $-\text{OCR}^{12}\text{R}^{13}\text{O}-$, form a 1,3-dioxolane ring;

where

R¹² and R¹³ are independently selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyalkyl, alkoxy, haloalkoxy, alkylthio, cyano, nitro, alkylsulfonyloxy, 20 alkylcarbonyl, alkoxycarbonyl, aryl, or aryloxy;

or

R¹² and R¹³ taken together with (=O), form 1,3-dioxol-2-one ring;

R^1 , R^2 , R^3 , R^4 , and R^5 are independently selected from hydrogen, halogen, alkyl, haloalkyl, hydroxyl, hydroxyalkyl, alkoxy, haloalkoxy, thio, alkylthio, haloalkylthio, pentahalothio, cyano, nitro, alkylsulfonyl, haloalkylsulfonyl, alkylsulfinyl, haloalkylsulfinyl, alkylsulfonyloxy, haloalkylsulfonyloxy, alkylcarbonyl, alkoxy carbonyl, dialkoxyalkylcarbonyl, alkoxy carbonylamino, alkylaminoxyalkyl, alkoxyiminoalkyl, alkenyloxyiminoalkyl, aryl, aryloxy, dioxanyl, dioxolanyl or either of R^1 and R^2 , or R^2 and R^3 , or R^3 and R^4 , or R^4 and R^5 taken together with $-OC(R^{19})_2O-$, $-OC(R^{19})_2(R^{19})_2O-$, $-OC(R^{19})_2(R^{19})_2-$, $-OC(R^{19})=N-$, or $-SC(R^{19})=N-$, forming a benzo-fused ring, where R^{19} is hydrogen, halogen, alkyl or haloalkyl; and,

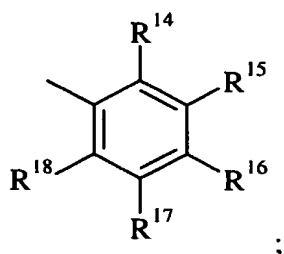
10 wherein at least one of R^1 , R^2 , R^3 , R^4 , and R^5 is other than hydrogen;

R^7 , R^8 , R^9 , R^{10} , and R^{11} are independently selected from hydrogen, halogen, alkyl, hydroxy, hydroxyalkyl, hydroxyalkoxy, alkoxy, alkoxyalkyl, alkoxyiminoalkyl, haloalkoxyiminoalkyl, cyanoalkoxyiminoalkyl, cyanoiminothioalkylamino, alkenyloxyiminoalkyl, alkynyloxyiminoalkyl, cycloalkoxy, cycloalkylalkoxy, phenoxy, alkoxy carbonylphenoxy, alkoxyalkoxy, alkoxyalkoxyalkoxy, alkylthio, alkylsulfonyl, alkylaminosulfonyl, dialkylaminosulfonyl, cycloalkylaminosulfonyl, alkenyloxy, alkynyloxy, haloalkenyloxy, alkylsulfonyloxy, optionally substituted arylalkoxy, cyano, nitro, amino, alkylamino, alkylcarbonylamino, alkoxy carbonylamino, cycloalkylalkoxy carbonylamino, alkenyloxy carbonylamino, alkynyloxy carbonylamino, haloalkylcarbonylamino, alkoxyalkoxy carbonylamino, alkoxy carbonylamino, alkoxy carbonyloxy, alkenylaminocarbonyloxy, alkynylaminocarbonyloxy, (alkyl)(alkoxy carbonyl)amino, alkylsulfonylamino, optionally substituted (heteroaryl)(alkoxy carbonyl)amino, optionally substituted arylcarbonylamino, alkoxy carbonyl, alkylaminocarbonyloxy, alkylaminocarbonylamino, dialkylaminocarbonylamino, alkylamino(thiocarbonyl)amino, dialkylphosphoroureidyl, acetoxyalkoxy, sulfonyloxyalkoxy, dialkoxyalkoxy, trialkoxyalkoxy, dialkoxyalkylacetal, trialkoxymethylorthoester, cyclic acetal, optionally substituted cyclic acetal, optionally substituted thienyl, optionally substituted 1,3-thiazolylalkoxy, optionally substituted aryl, optionally substituted aryloxy, optionally substituted aryloxyalkyl, optionally substituted arylaminocarbonyloxy, optionally substituted

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arylalkoxycarbonylamino, optionally substituted heteroaryl, optionally substituted
 heteroaryloxy, optionally substituted pyrrolyl, optionally substituted pyrazolyl,
 optionally substituted pyrazinyloxy, optionally substituted cycloalkylcarbonylamino,
 optionally substituted 1,3-oxazolinyl, optionally substituted 1,3-oxazolinyl, 5
 optionally substituted 1,3-oxazolinylamino, optionally substituted 1,2,4-triazolyl,
 optionally substituted 1,2,3-thiadiazolyl, optionally substituted 1,2,5-thiadiazolyl,
 optionally substituted 1,2,5-thiadiazolyloxy, optionally substituted 2H-tetrazolyl,
 optionally substituted pyridyl, optionally substituted pyridyloxy, optionally
 substituted pyridylamino, optionally substituted pyrimidinyl, optionally substituted
 10 pyrimidinyl, optionally substituted 3,4,5,6-tetrahydropyrimidinyl, optionally
 substituted pyridazinyloxy, or optionally substituted 1,2,3,4-tetrahydronaphthalenyl,
 wherein the optional substituent is selected from one or more of halogen, alkyl,
 haloalkyl, alkoxy, cyano, nitro, amino, alkylcarbonyl, alkoxycarbonyl,
 alkoxyiminoalkyl, dialkylacetal, alkylthiol, alkylsulfoxide, or alkoxycarbonylamino;
 15 and, wherein at least one of R^7 , R^8 , R^9 , R^{10} , and R^{11} is other than hydrogen;

R is alkyl, cycloalkyl, alkenyl, alkoxycarbonyl, optionally substituted pyrid-2-yl
 wherein the optional substituent is selected from hydrogen, halogen, haloalkoxy or
 haloalkyl,
 20 or
 substituted phenyl have the following structure,



where
 25 R^{14} , R^{15} , R^{16} , R^{17} , and R^{18} are independently selected from hydrogen, halogen, alkyl,
 haloalkyl, hydroxyl, hydroxyalkyl, alkoxy, haloalkoxy, thio, alkylthio, haloalkylthio,
 pentahalothio, cyano, nitro, alkylsulfonyl, haloalkylsulfonyl, alkylsulfinyl,
 haloalkylsulfinyl, alkylsulfonyloxy, haloalkylsulfonyloxy, alkylcarbonyl,

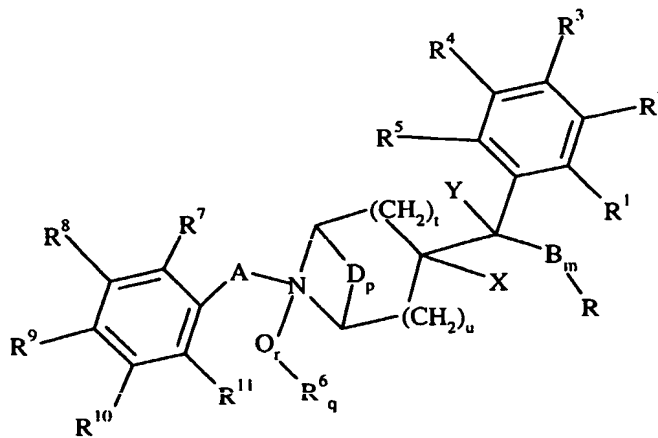
alkoxycarbonyl, dialkoxyalkylcarbonyl, alkoxycarbonylamino, alkylaminoxyalkyl, alkoxyiminoalkyl, alkenyloxyiminoalkyl, aryl, aryloxy, dioxanyl, dioxolanyl or either of R^{14} and R^{15} , or R^{15} and R^{16} , or R^{16} and R^{17} , or R^{17} and R^{18} taken together with $-\text{OC}(R^{19})_2\text{O}-$, $-\text{OC}(R^{19})_2(R^{19})_2\text{O}-$, $-\text{OC}(R^{19})_2(R^{19})_2-$, $-\text{OC}(R^{19})=\text{N}-$, or $-\text{SC}(R^{19})=\text{N}-$, forming a benzo-fused ring, where R^{19} is hydrogen, halogen, alkyl or haloalkyl; and, wherein at least one of R^{14} , R^{15} , R^{16} , R^{17} , and R^{18} is other than hydrogen;

A is selected from $-\text{CH}_2-$, $-\text{CH}_2\text{CH}_2-$, $-\text{CH}_2\text{CH}_2\text{CH}_2-$, $-\text{OCH}_2\text{CH}_2-$, $-\text{OCH}_2\text{CH}_2\text{CH}_2-$, $-\text{OCH}_2\text{CH}_2\text{CH}_2\text{CH}_2-$, $-\text{OCH}_2\text{CH}(\text{OH})\text{CH}_2-$, $-\text{NHCH}_2\text{CH}_2-$, $-\text{N}(\text{CH}_3)\text{CH}_2\text{CH}_2-$, $-\text{N}[\text{C}(=\text{O})\text{CH}_3]\text{CH}_2\text{CH}_2-$, or $-\text{N}[\text{C}(=\text{O})\text{OCH}_3]\text{CH}_2\text{CH}_2-$;

B is selected from $-\text{O}-$, $-\text{S}-$, $-\text{CH}_2\text{O}-$, $-\text{OCH}_2-$, $-\text{OC}(=\text{O})\text{NH}-$, $-\text{OC}(=\text{O})\text{O}-$, or $-\text{NHSO}_2-$;

R^6 is selected from alkyl, haloalkyl, hydroxyalkyl, alkoxyalkyl, dialkylaminoalkyl, alkylaminocarbonyloxyalkyl, alkylthioalkyl, alkylsulfonylalkyl, alkylcarbonyloxyalkyl, alkoxycarbonylalkyl, carboxyalkyl, arylalkyl, arylcarbonyl, sulfonato, or sulfonatoalkyl, and may bear a negative charge resulting in an inner salt; and a separate ion is chloride, bromide, iodide, or an alkyl or phenyl sulfate or sulfonate.

Another embodiment of the present invention is a compound of formula I:



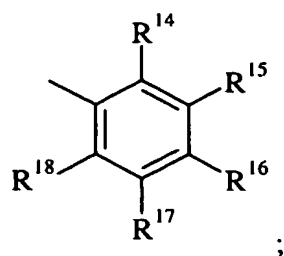
I

wherein;

- 5 r is selected from 0 or 1; m, q and p are 0; t and u are 1;
 A is $-\text{CH}_2-$;
 X is selected from halogen or hydroxyl;
 Y is selected from hydrogen or hydroxyl;
 R^1 , R^2 , R^3 and R^4 are independently selected from hydrogen, halogen, alkyl, alkoxy,
 10 haloalkyl, haloalkoxy or $-\text{CH}=\text{NOC}_2\text{H}_5$;
 R^5 is hydrogen;
 R^7 , R^8 , R^{10} and R^{11} are hydrogen;
 R^9 is selected from $-\text{OC}_2\text{H}_5$, $-\text{OC}_3\text{H}_7$, $-\text{OCH}(\text{CH}_3)_2$, $-\text{OCH}_2\text{CH}_2\text{OCH}_3$, $-\text{OCH}_2\text{CH}_2\text{CH}_2\text{OCH}_3$, cyclopropylmethoxy, 2-chlorophenoxy, 3-chlorophenoxy, 4-
 15 chlorophenoxy, pyrimidin-2-yl, pyrid-2-yl, 3-chloro-pyrid-2-yl, 3-methyl-pyrid-2-
 yloxy, 4-methyl-pyrid-2-yloxy, 5-methyl-pyrid-2-yloxy, 6-methyl-pyrid-2-yloxy, 3-
 chloro-pyrid-2-yloxy, 3-trifluoromethyl-pyrid-2-yloxy, 3-cyano-pyrid-2-yloxy, 5-
 cyano-pyrid-2-yloxy, 6-dimethoxymethyl-pyrid-2-yloxy, pyrid-2-yloxy,
 $\text{CO}_2\text{CH}(\text{CH}_3)_2$, $-\text{CH}=\text{NOCH}_3$, $-\text{CH}=\text{NOC}_2\text{H}_5$, $-\text{CH}=\text{NOCH}_2\text{CF}_3$, $-\text{CH}=\text{NOallyl}$, $-\text{CH}=\text{NOCH}_2\text{CH}=\text{CH}_2$, $-\text{CH}=\text{NOCH}_2\text{CN}$, $-\text{CH}=\text{NOCH}(\text{CH}_3)_2$, $-\text{CH}=\text{NOCH}_2\text{C}\equiv\text{CH}$, $-\text{CH}=\text{NOCH}_2\text{CH}_2\text{F}$, $-\text{CH}=\text{NOCH}_2\text{CH}_2\text{OCH}_3$, $-\text{CH}=\text{NOCH}_2\text{OC}_2\text{H}_5$, $-\text{CH}=\text{NOCH}_2\text{CH}_2\text{OCH}_2\text{CH}_2\text{OCH}_3$, $-\text{NHCO}_2\text{CH}_3$, $-\text{NHCO}_2\text{C}_2\text{H}_5$, $-\text{NHCO}_2\text{CH}(\text{CH}_3)_2$, $-\text{NHCO}_2\text{CH}_2-c-\text{C}_3\text{H}_5$, $-\text{CH}(\text{OH})\text{C}_6\text{H}_5-p\text{-Cl}$, $-\text{OC}(=\text{O})\text{NHCH}_3$, $-\text{OC}(=\text{O})\text{NHC}_2\text{H}_5$, $-\text{OC}(=\text{O})\text{NHCH}(\text{CH}_3)_2$, $-\text{NHC}(\text{SCH}_3)=\text{NCN}$, pyrimidin-2-yloxy, 6-chloro-pyridazin-
 20 3yloxy, 6-methoxy-pyridazin-3yloxy, 6-methyl-pyridazin-3yloxy, 2-methyl-2H-
 tetrazol-5-yl, 2-ethyl-2H-tetrazol-5-yl, 1,3-dioxan-2-yl or 5,5-dimethyl-1,3-dioxan-
 2-yl; and

R is phenyl substituted with R^{14} , R^{15} , R^{16} , R^{17} , and R^{18} ,

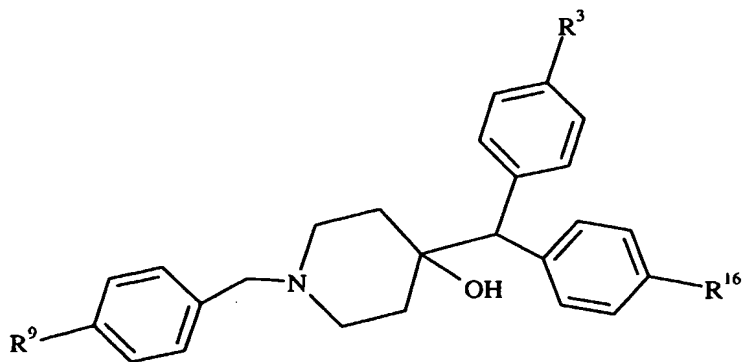
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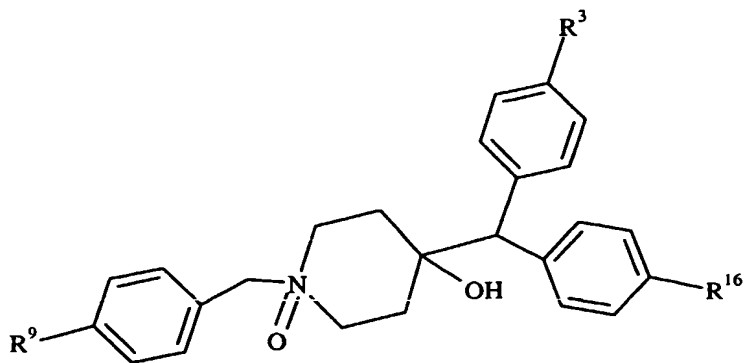
where

R^{16} is selected from haloalkyl or haloalkoxy, and R^{14} , R^{15} , R^{17} and R^{18} are hydrogen.

5 Another embodiment of the present invention is a compound of formula **I-H** or **I-J**:



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I-H**I-J**

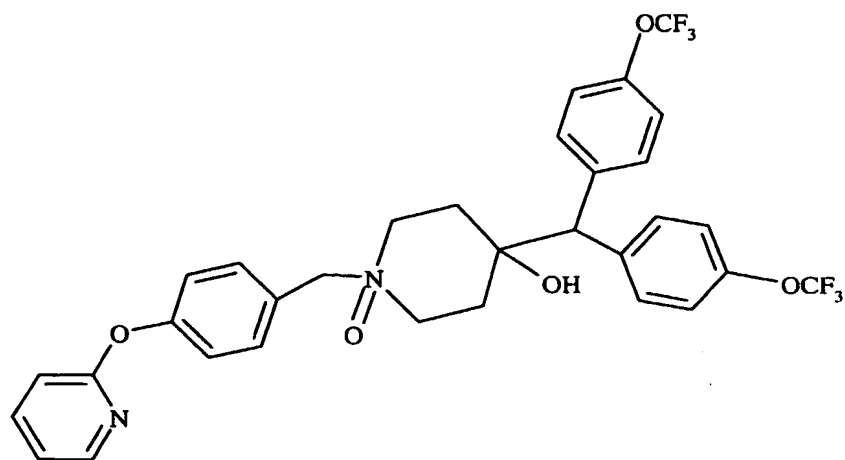
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wherein,

R³ is haloalkyl or haloalkoxy;

R⁹ is selected from -OCH₂CH₂OCH₃, pyrid-2-yloxy, pyrid-2-yl, 3-cyano-pyrid-2-yloxy, 5-methyl-pyrid-2-yloxy, pyrimidin-2-yloxy, pyrimidin-2-yl, 6-chloro-pyridazin-3-yloxy or 6-methoxy-pyridazin-3-yloxy; and R¹⁶ is haloalkyl or haloalkoxy.

Yet another embodiment of the present invention is the compound:



namely, 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-4-hydroxy-1-[(4-(2-pyridyloxy)phenyl)methyl]piperidin-1-oxide, and agriculturally-acceptable salts thereof.

In certain cases the compounds within the scope of formula I may possess asymmetric centers, which can give rise to optical enantiomorphs and diastereomers. Compounds within the scope of formula I may exist in two or more forms, i.e., polymorphs, which are significantly different in physical and chemical properties. Compounds within the scope of formula I may also exist as tautomers, which are in equilibrium. Compounds within the scope of formula I may also possess acidic or basic moieties, which may allow for the formation of agriculturally acceptable salts or agriculturally acceptable metal complexes.

This invention includes the use of such enantiomorphs, polymorphs, tautomers, salts and metal complexes. Agriculturally acceptable salts and metal complexes include, without limitation, for example, ammonium salts, the salts of

organic and inorganic acids, such as hydrochloric acid, oleic acid, octanoic acid, 2-ethylhexanoic acid, alkyl sulfonic acid, ethanesulfonic acid, trifluoroacetic acid, methylbenzenesulfonic acid, phosphoric acid, gluconic acid, pamoic acid, and other acid salts, and the alkali metal and alkaline earth metal complexes with, for example, sodium, potassium, lithium, magnesium, calcium, and other metals.

The methods of the present invention are predicated on causing an insecticidally effective amount of a compound of formula I to be present within insects in order to kill or control the insects. Preferred insecticidally effective amounts are those that are sufficient to kill the insect. It is within the scope of the present invention to cause a compound of formula I to be present within insects by contacting the insects with a derivative of that compound, which derivative is converted within the insect to a compound of formula I. This invention includes the use of such compounds, which can be referred to as pro-insecticides.

Another aspect of the present invention relates to compositions containing an insecticidally effective amount of at least one compound of formula I, and, optionally, an effective amount of at least one second compound, with at least one agriculturally acceptable extender or adjuvant.

Another aspect of the present invention relates to methods of controlling insects by applying an insecticidally effective amount of a composition set forth above to a locus of crops such as, without limitation, cereals, cotton, vegetables, and fruits, other areas where insects are present or are expected to be present, or adjacent to areas where insects are present or are expected to be present.

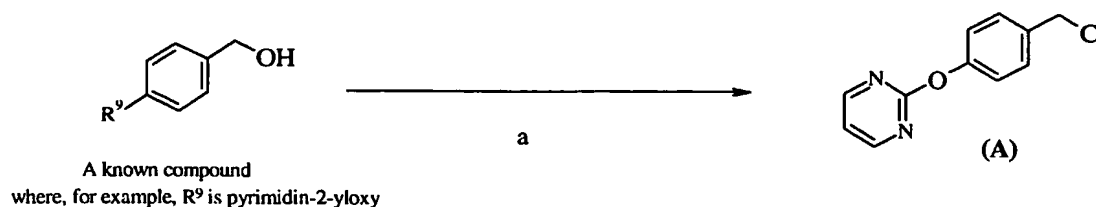
The present invention also includes the use of the compounds and compositions set forth herein for control of insects in greenhouse crops, nursery crops, ornamentals, turfs, forestry, stored food and fiber products, structures, livestock, households, and public and animal health, for example, ants, flies, cockroaches, white grubs, dry wood termites and subterranean termites as well as other insects; and also for use in promotion of animal and human health as pharmaceutical agents and compositions thereof.

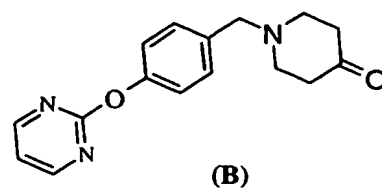
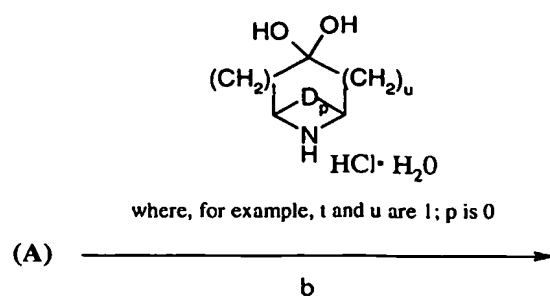
As used in this specification and unless otherwise indicated the substituent terms "alkyl", "alkenyl", "alkynyl", "alkoxy", "alkenyloxy", and "alkynyloxy" used alone or as part of a larger moiety, includes straight or branched chains of at least

one or two carbon atoms, as appropriate to the substituent, and preferably up to 12 carbon atoms, more preferably up to ten carbon atoms, most preferably up to seven carbon atoms, wherein "alkenyl" has at least one carbon to carbon double bond, and "alkynyl" has at least one carbon to carbon triple bond. The term "aryl" refers to an aromatic ring structure, including fused rings, having four to ten carbon atoms, for example, phenyl and naphthyl. The term "heteroaryl" refers to an aromatic ring structure, including fused rings, having four to ten carbon atoms, and in which one or more of the atoms in the ring is other than carbon, for example, sulfur, oxygen, or nitrogen. The term "THF" refers to tetrahydrofuran. The term "DMSO" refers to methyl sulfoxide. The term "DMF" refers to N,N-dimethylformamide. The term "halogen" or "halo" refers to fluorine, bromine, iodine, or chlorine. The term "ambient temperature" or "room temperature" often abbreviated as "RT", for example, in reference to a chemical reaction mixture temperature, refers to a temperature in the range of 20 °C to 30 °C.

Scheme 1 below illustrates a general procedure for synthesizing compounds of formula I, where, for example, m, p, and q are 0; t and u are 1; r is 0 or 1, and if r is 1 an N-oxide is formed; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷ and R¹⁸; X is OH or F; Y, R¹, R², R⁴, R⁵, R¹⁴, R¹⁵, R¹⁷, R¹⁸, R⁷, R⁸, R¹⁰ and R¹¹ are H; R³ and R¹⁶ are -OCF₃; and R⁹ is pyrimidin-2-yloxy.

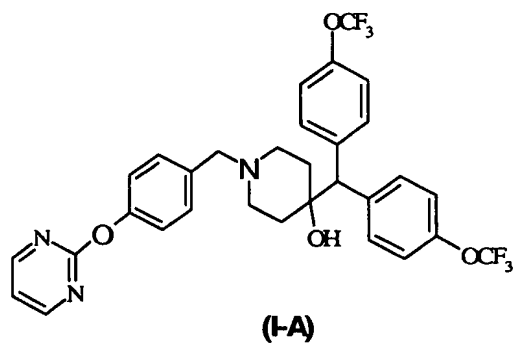
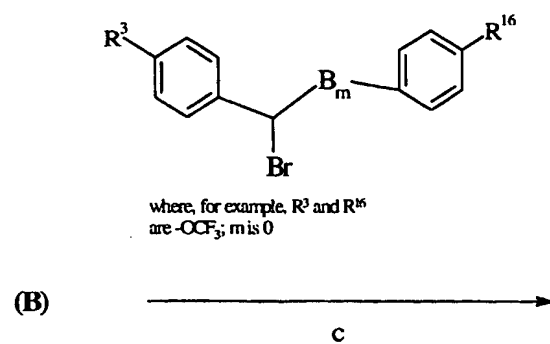
Scheme 1

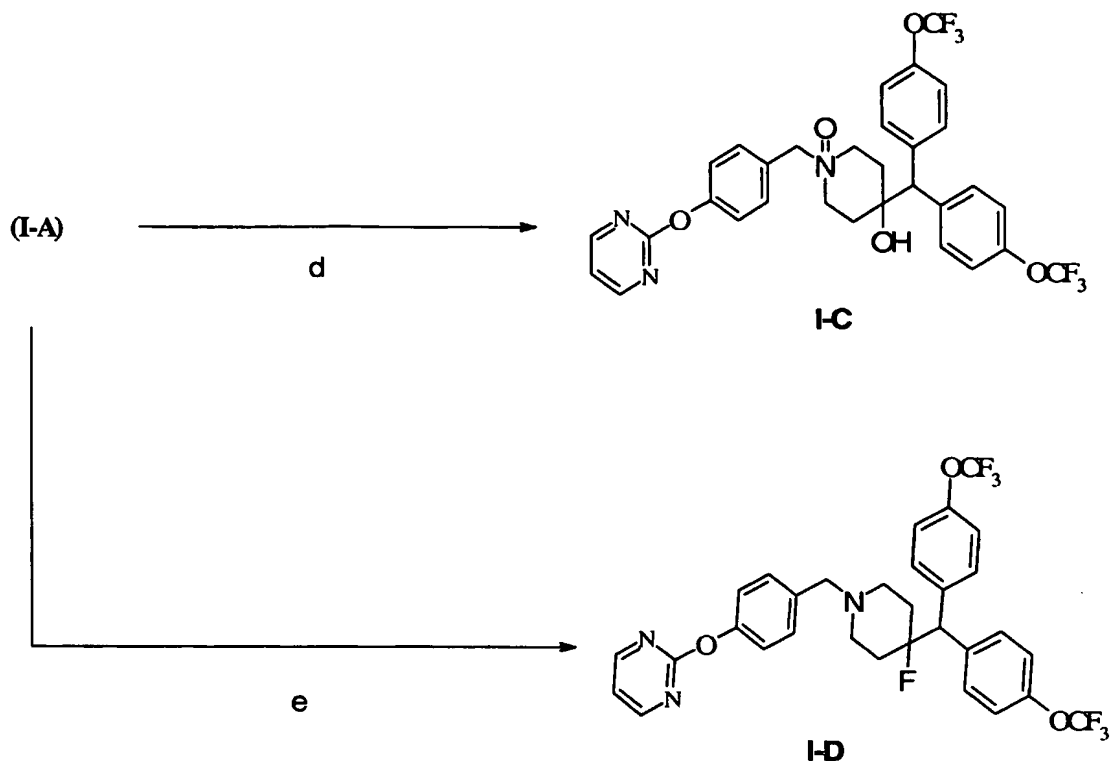




Scheme 1 (cont'd)

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a) $\text{SOCl}_2/\text{C}_5\text{H}_5\text{N}/\text{CH}_2\text{Cl}_2/15^\circ\text{C}$ b) $(\text{CH}_3)_2\text{SO}/\text{C}_8\text{H}_{19}\text{N}/\text{RT}$ c) BuLi (2.5M)/THF/-78 °C
 d) $\text{H}_2\text{O}_2/\text{CH}_3\text{OH}/\text{RT}$ e) $\text{C}_2\text{H}_5\text{NSF}_3/\text{CH}_2\text{Cl}_2/-40^\circ\text{C}$

5

In a first step as depicted in Scheme 1, an appropriately substituted alcohol, for example, the known compound (4-pyrimidin-2-yloxyphenyl)methan-1-ol, is halogenated with, for example thionyl chloride, to afford the corresponding 2-[4-(chloromethyl)phenoxy]pyrimidine (A). Intermediate (A) is then reacted under

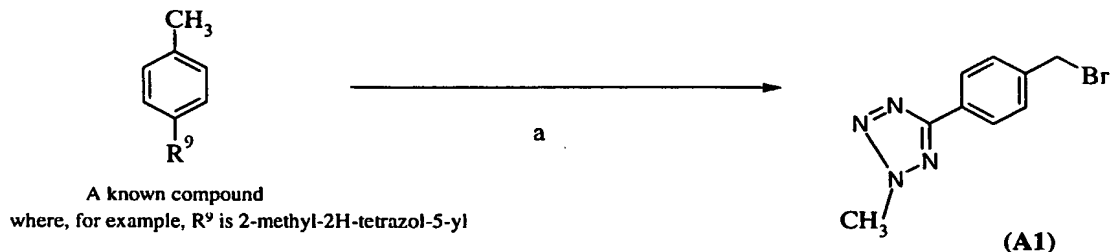
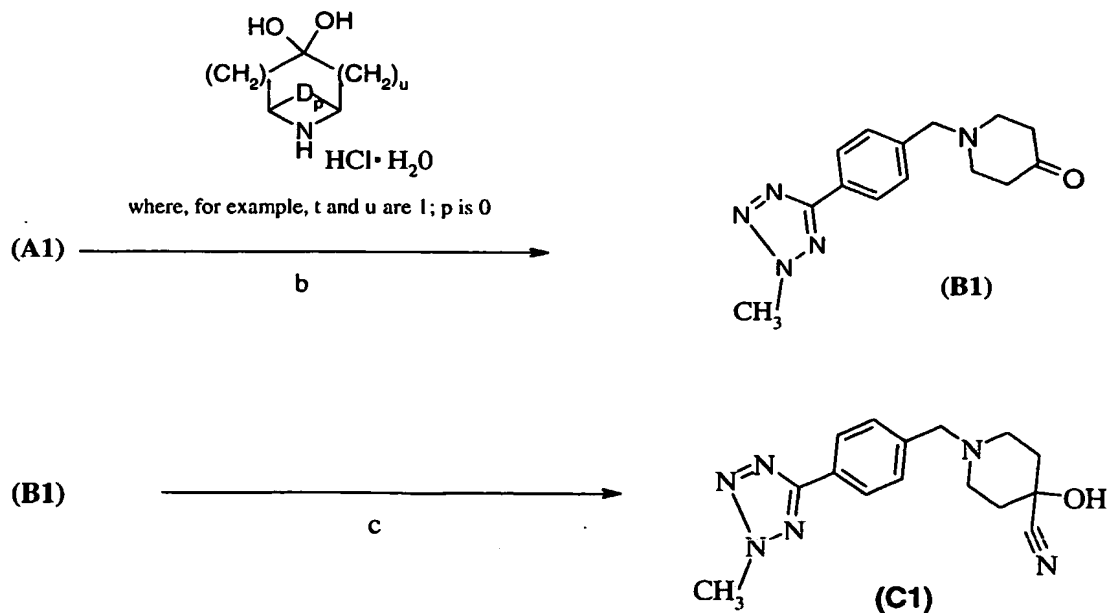
10 basic conditions with an appropriately substituted cyclic amine derivative, for example, the known compound 4-piperidone hydrochloride monohydrate, to afford the corresponding 1-[(4-pyrimidin-2-yloxyphenyl)methyl]piperidin-4-one (B). A mixture of intermediate (B) and an appropriately substituted haloalkyl derivative, for example bis(trifluoromethoxyphenyl)bromomethane, is reacted in the presence of n-

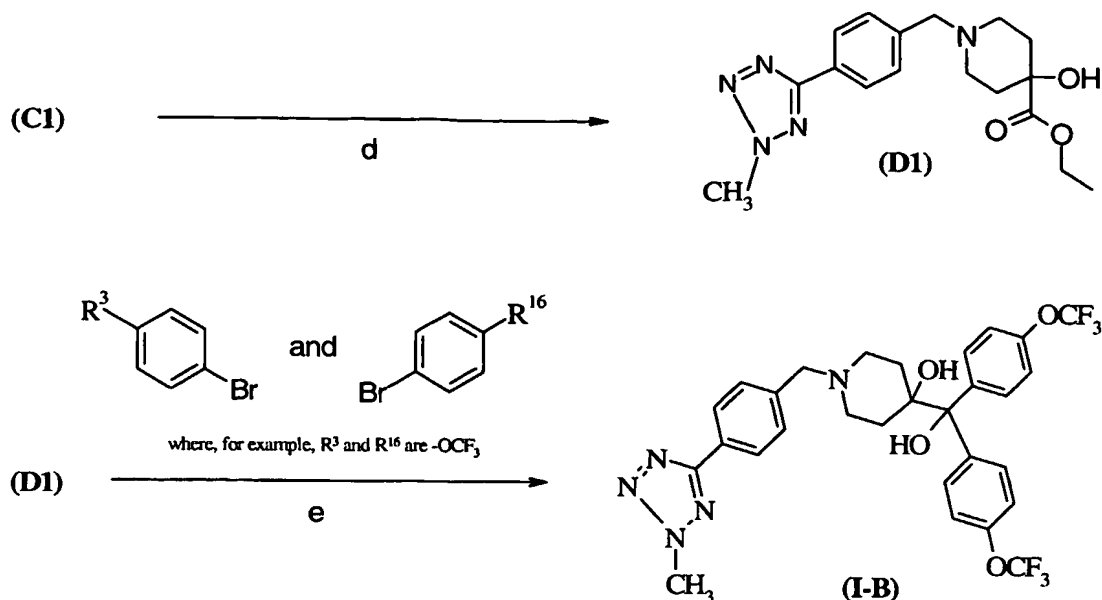
15 butyl lithium, to afford the corresponding 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-1-[(4-pyrimidin-2-yloxyphenyl)methyl]piperidin-4-ol (I-A). Intermediate (I-A) is then oxidized, with for example hydrogen peroxide, in an appropriate solvent, to form an N-oxide, a compound of formula I-C. In a separate synthesis, intermediate (I-A) is reacted with a thiohalide, for example

(dimethylamino)sulfur trifluoride, to provide halogen-derived compounds of formula I-D, wherein X is, for example fluorine. Examples 1 and 3, set forth below, provide detailed methods to how compounds of formula I (-A, -C and -D) shown in Scheme 1 were prepared.

5

Scheme 2 below illustrates a general procedure for synthesizing compounds of formula I similar to those set forth in Scheme 1, differing in that Y is OH; X is OH; r is 0; and R⁹ is 2-methyl-2H-tetrazol-5-yl.

10 Scheme 215 Scheme 2 (cont.)



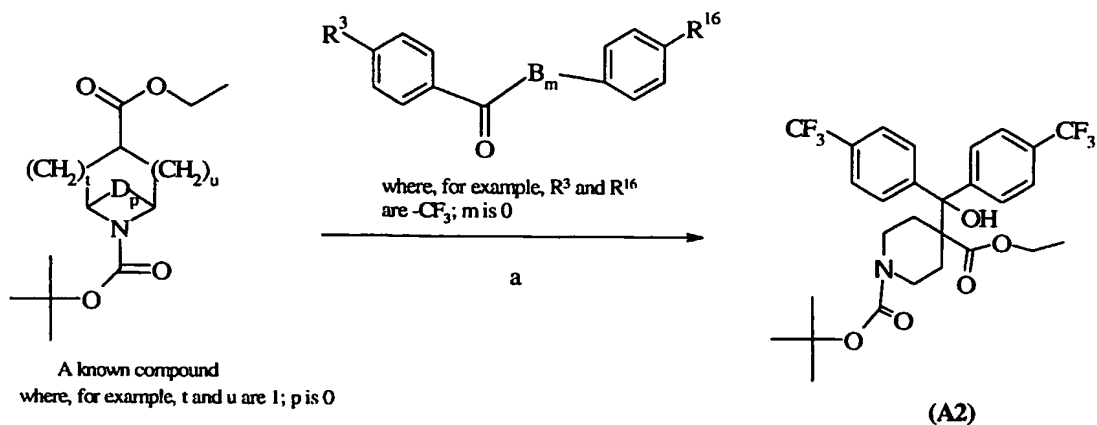
- 5 a) $C_4H_4O_2NBr/CCl_4/(C_6H_5CO)_2O_2/light$ b) $[(CH_3)_2CH]_2NC_2H_5/(CH_3)_2SO/RT$
 c) $NaCN/HCl/H_2O/(C_2H_5)_2O/10\text{ }^\circ C$ d) $HCl/C_2H_5OH/55\text{ }^\circ C$ e) $Mg/THF/50\text{ }^\circ C$

In a first step as depicted in Scheme 2, an appropriately substituted phenyl derivative, for example, the known compound 2-methyl-5-(4-methylphenyl)-1,2,3,4-tetrazole, is brominated with, for example N-bromosuccinimide and light, to afford the corresponding 5-[4-(bromomethyl)phenyl]-2-methyl-1,2,3,4-tetrazole (A1). Intermediate (A1) is then reacted under basic conditions with an appropriately substituted cyclic amine derivative, for example, the known compound 4-piperidone hydrochloride monohydrate, to afford the corresponding 1-{[4-(2-methyl-1,2,3,4-tetrazol-5-yl)phenyl]methyl}piperidin-4-one (B1). Intermediate (B1) is then reacted under acidic conditions with, for example sodium cyanide, to afford the corresponding nitrile compound, 4-hydroxy-1-{[4-(2-methyl(1,2,3,4-tetrazol-5-yl))phenyl]methyl}piperidine-4-carbonitrile (C1). Intermediate (C1) is esterified under acidic conditions with, for example ethanol, to afford the corresponding ethyl 4-hydroxy-1-{[4-(2-methyl-(1,2,3,4-tetrazol-5-yl))phenyl]methyl}piperidine-4-carboxylate (D1). The Grignard of an appropriately substituted halophenyl derivative, for example 4-trifluoromethoxybromobenzene, is reacted with intermediate (D1) to provide compounds of formula I-B, wherein X and Y are, for

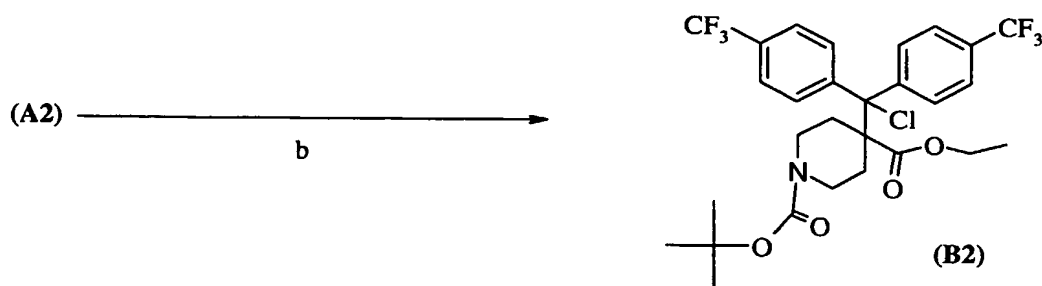
example hydroxyl. Example 2, set forth below, provides detailed methods to how compounds of formula I (-B) shown in Scheme 2 were prepared.

Scheme 3 below illustrates a general procedure for synthesizing compounds of formula I similar to those set forth in Scheme 1, differing in that Y is Cl; X is -CO₂C₂H₅; r is 0; R³ and R¹⁶ are -CF₃; and R⁹ is pyrid-2-yloxy.

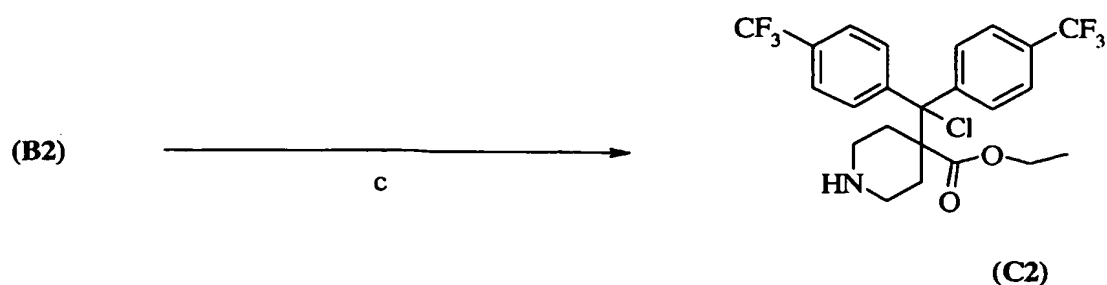
Scheme 3



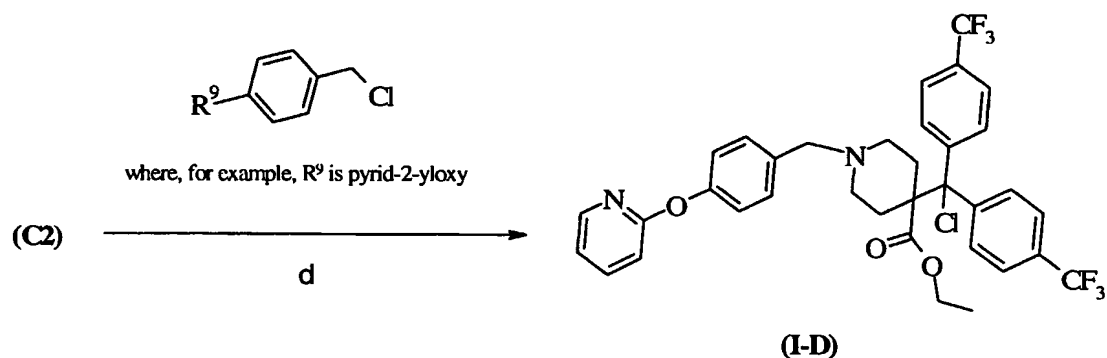
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Scheme 3 (cont'd)

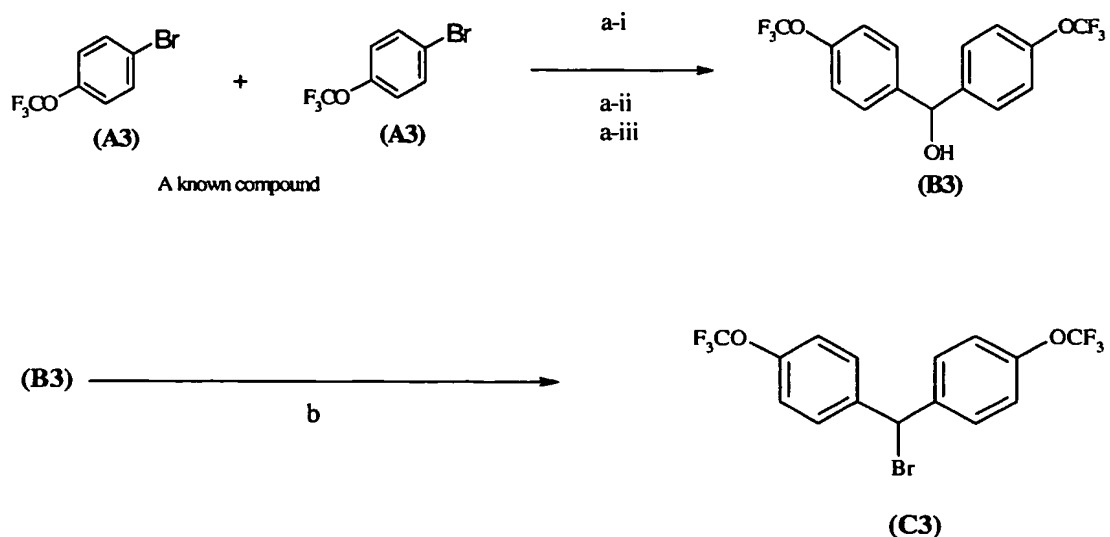
- 10 a) BuLi (2.5 M)/THF/[(CH₃)₂CH]₂NH/-78 °C b) SOCl₂/C₅H₅N/CH₂Cl₂/RT
c) CF₃CO₂H/CH₂Cl₂/RT d) C₈H₁₉N/DMF/RT

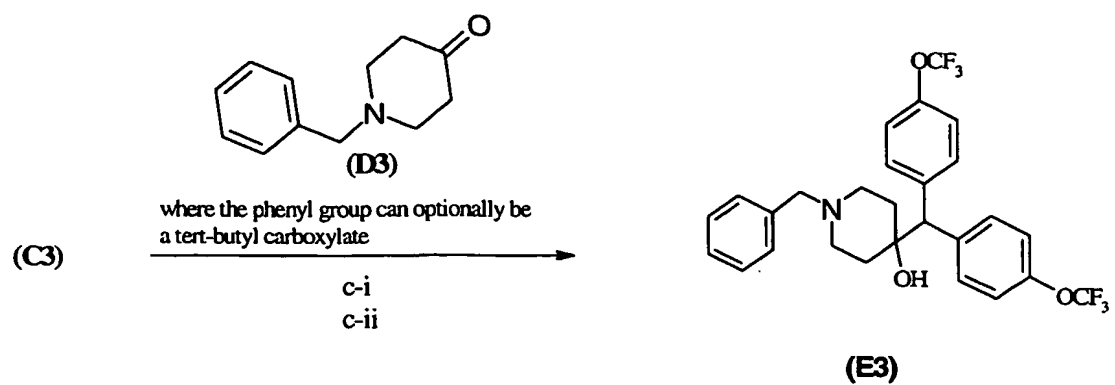
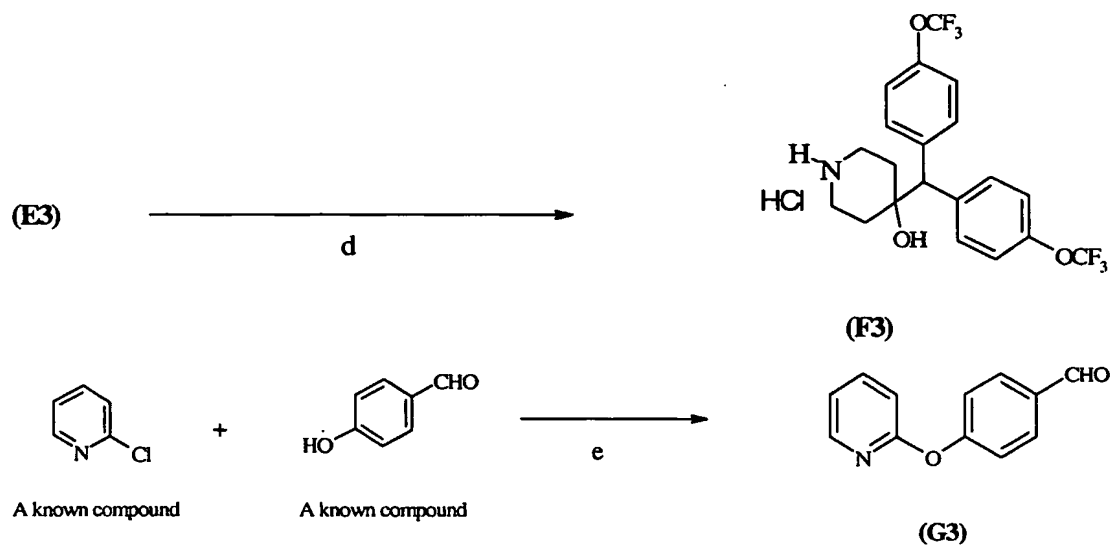
In a first step as depicted in Scheme 3, an appropriately substituted piperidine, for example, the known compound ethyl 1-[(tert-butyl)oxycarbonyl]piperidine-4-carboxylate, is enolated, with for example lithium diisopropylamide, and then reacted under basic conditions with an appropriately substituted phenyl ketone, for example di-4-(trifluoromethyl)phenyl ketone, to afford the corresponding ethyl 1-[(tert-butyl)oxycarbonyl]-4-{bis[4-(trifluoromethyl)phenyl]hydroxymethyl}piperidine-4-carboxylate (A2). Intermediate (A2) is then chlorinated with, for example thionyl chloride, to afford the corresponding ethyl 1-[(tert-butyl)oxycarbonyl]-4-{bis[4-(trifluoromethyl)phenyl]chloromethyl}piperidine-4-carboxylate (B2). The (tert-butyl)oxycarbonyl group is cleaved under acidic conditions from the piperidine ring

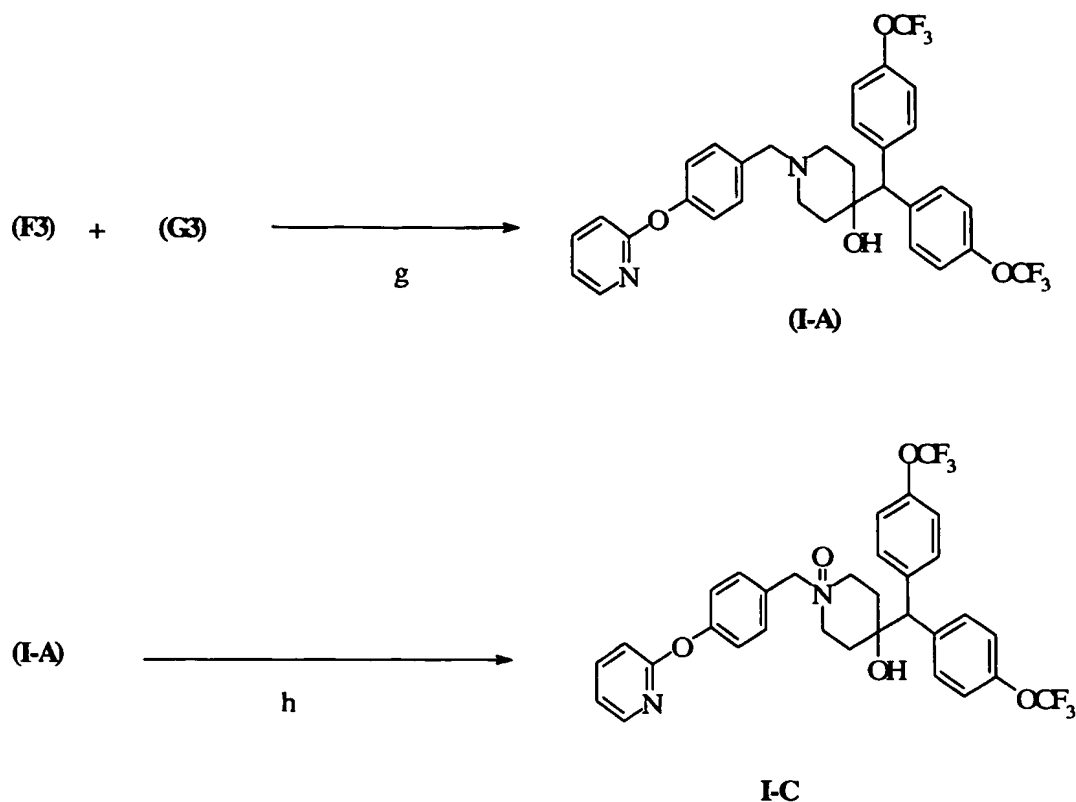
of (B2), to afford the corresponding 4-{bis[4-(trifluoromethyl)phenyl]chloromethyl}piperidine-4-carboxylate (C2). Intermediate (C2) is then reacted under basic conditions with an appropriately substituted haloalkylphenyl derivative, for example 2-[4-(chloromethyl)phenoxy]pyridine, to provide compounds of formula I-D, wherein X is, for example an alkoxycarbonyl and Y is, for example chlorine. Example 4, set forth below, provides detailed methods to how compounds of formula I (-D) shown in Scheme 3 were prepared.

Scheme 4 below illustrates a general procedure for synthesizing compounds of formula I, where, for example, m, p, and q are 0; t and u are 1; r is 0 or 1, and if r is 1 an N-oxide is formed; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷ and R¹⁸; X is OH; Y, R¹, R², R⁴, R⁵, R¹⁴, R¹⁵, R¹⁷, R¹⁸, R⁷, R⁸, R¹⁰ and R¹¹ are H; R³ and R¹⁶ are -OCF₃; and R⁹ is pyrid-2-yloxy.

15 Scheme 4



5 Scheme 4 (cont'd)



5 a-i) THF/I-PrMgCl/15°C to RT/24 hours a-ii)HCO₂Et/-10°C to 0°C a-iii)10% NH₄Cl b)
 Heptane/HBr/Acetic Acid/RT/3 hours c-i) t-BuLi /THF/-85°C to -60°C/12 hours c-ii) HCl(g) d)
 MeOH/HCOOH/Pd(OH)₂/C'/40°C to 55°C/4 hours e) K₂CO₃/Cu₂O/145°C to 170°C/3.5 hours g)
 THF/NaBH(OAc)₃/RT/12 hours h)50%H₂O₂/MeOH/40°C to 55°C /9-44 hours

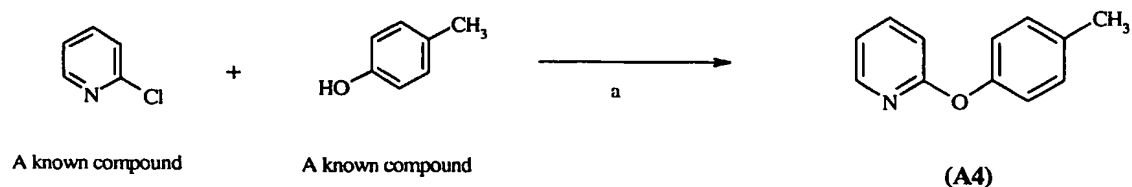
In the first step as depicted in Scheme 4, two appropriately substituted aryl
 halides, for example, the known compound 4-bromo-1-(trifluoromethoxy)benzene
 10 (A3), were cross-coupled with a Grignard reagent and an alkyl formate, for example,
 ethyl formate to form bis[4-(trifluoromethoxy)phenyl]methan-1-ol (B3).
 Intermediate (B3) was then reacted under acidic conditions with hydrogen bromide,
 to afford the corresponding bis(trifluoromethoxyphenyl)bromomethane (C3).
 Intermediate (C3) was then lithiated, for example with butyl lithium, and then
 15 reacted with an appropriately N-substituted piperidin-4-one, formula (D3), for
 example 1-benzylpiperidin-4-one, at a temperature in the range of -85°C to -60°C to
 afford the corresponding 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-1-
 benzylpiperidin-4-ol (E3). Intermediate (E3) was then reacted with an acid, for
 example formic acid, in the presence of a catalyst, for example a palladium catalyst,

to form the hydrogen chloride salt of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}piperidin-4-ol (**F3**). Next, an appropriately substituted phenol, for example, the known compound 4-hydroxybenzaldehyde, was cross-coupled with a halopyridine, for example 2-chloropyridine, in the presence of potassium carbonate and a catalytic amount of copper oxide at a temperature in the range of 145°C to 170°C to form 4-(2-pyridyloxy)benzaldehyde (**G3**). Intermediate (**F3**) was then cross-coupled with Intermediate (**G3**) in the presence of sodium triacetoxymethylborohydride to form 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-1-[(4-(2-pyridyloxy)phenyl)methyl]piperidin-4-ol (**I-A**). Intermediate (**I-A**) was then oxidized with hydrogen peroxide at a temperature in the range of 40°C to 55°C to form a compound of formula **I-C**.

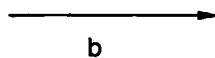
Scheme 5 below illustrates a general procedure for synthesizing compounds of formula **I**, where, for example, m, p, and q are 0; t and u are 1; r is 0 or 1, and if r is 1 an N-oxide is formed; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷ and R¹⁸; X is OH; Y, R¹, R², R⁴, R⁵, R¹⁴, R¹⁵, R¹⁷, R¹⁸, R⁷, R⁸, R¹⁰ and R¹¹ are H; R³ and R¹⁶ are -OCF₃; and R⁹ is pyrid-2-yloxy.

Scheme 5

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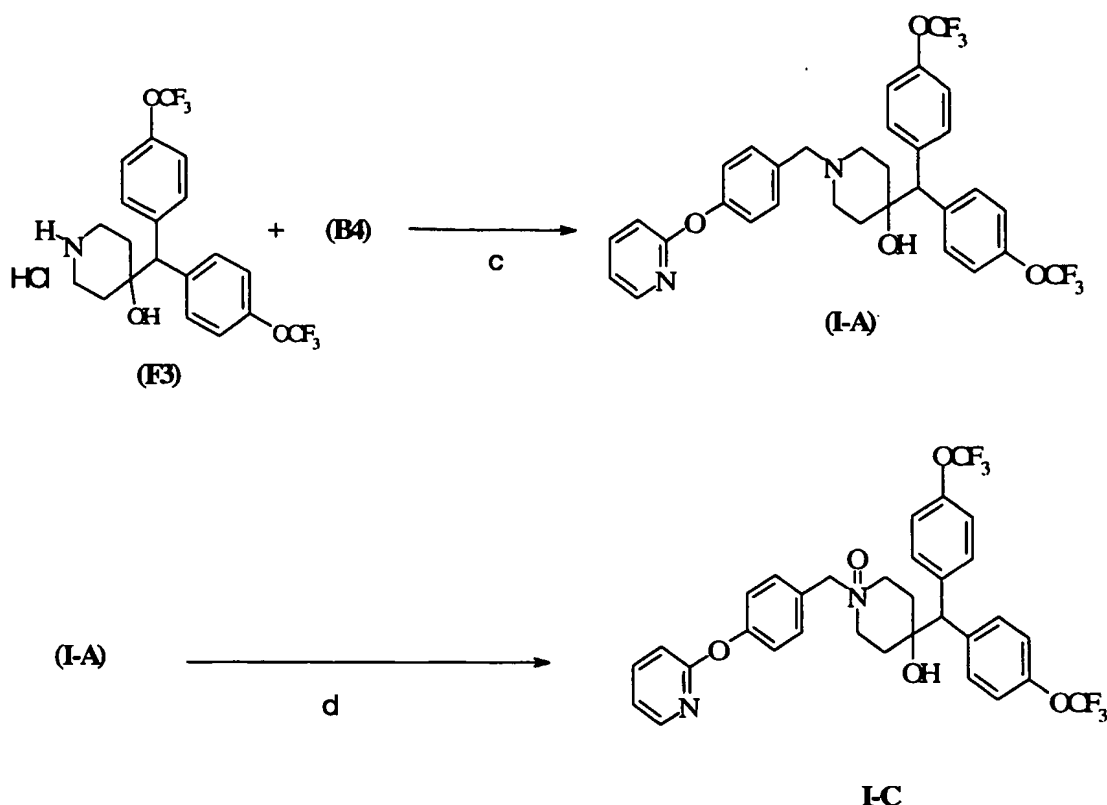


(A4)



(B4)

25



a) $K_2CO_3/Cu_2O/145^\circ C$ to $170^\circ C/3.5$ hours b) Br_2 c) K_2CO_3/DMF d) $50\%H_2O_2/MeOH/40^\circ C$ to $55^\circ C/9-44$ hours

- 5 In the first step of Scheme 5, an appropriately substituted phenol, for example, the known compound 4-methyl phenol, can be cross-coupled with a halopyridine, for example 2-chloropyridine, in the presence of potassium carbonate and a catalytic amount of copper oxide at a temperature in the range of $145^\circ C$ to $170^\circ C$ to form 2-(4-methylphenoxy)pyridine (A4). Intermediate (A4) can then be
- 10 halogenated with, for example bromine, to form 2-[4-(bromomethyl)phenoxy]pyridine (B4). Intermediate (F3), made as in Scheme 4, can then be cross-coupled with Intermediate (B4) in the presence of potassium carbonate to form 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-1-[(4-(2-pyridyloxy)phenyl)methyl]piperidin-4-ol (I-A). Intermediate (I-A) can then be
- 15 oxidized as in Scheme 4 to form a compound of formula I-C.

One skilled in the art will, of course, recognize that the formulation and mode of application of a toxicant may affect the activity of the material in a given application. Thus, for agricultural use the present insecticidal compounds may be

formulated as a granular product of relatively large particle size (for example, 8/16 or 4/8 US Mesh), as water-soluble or water-dispersible granules, as powdery dusts, as wettable powders, as emulsifiable concentrates, as aqueous emulsions, as solutions, or as any of other known types of agriculturally-useful formulations, depending on the desired mode of application. It is to be understood that the amounts specified in this specification are intended to be approximate only, as if the word "about" were placed in front of the amounts specified.

These insecticidal compositions may be applied either as water-diluted sprays, or dusts, or granules to the areas in which suppression of insects is desired. These formulations may contain as little as 0.1%, 0.2% or 0.5% to as much as 95% or more by weight of active ingredient.

Dusts are free flowing admixtures of the active ingredient with finely divided solids such as talc, natural clays, kieselguhr, flours such as walnut shell and cottonseed flours, and other organic and inorganic solids which act as dispersants and carriers for the toxicant; these finely divided solids have an average particle size of less than about 50 microns. A typical dust formulation useful herein is one containing 1.0 part or less of the insecticidal compound and 99.0 parts of talc.

Wettable powders, also useful formulations for insecticides, are in the form of finely divided particles that disperse readily in water or other dispersant. The wettable powder is ultimately applied to the locus where insect control is needed either as a dry dust or as an emulsion in water or other liquid. Typical carriers for wettable powders include Fuller's earth, kaolin clays, silicas, and other highly absorbent, readily wet inorganic diluents. Wettable powders normally are prepared to contain about 5-80% of active ingredient, depending on the absorbency of the carrier, and usually also contain a small amount of a wetting, dispersing or emulsifying agent to facilitate dispersion. For example, a useful wettable powder formulation contains 80.0 parts of the insecticidal compound, 17.9 parts of Palmetto clay, and 1.0 part of sodium lignosulfonate and 0.3 part of sulfonated aliphatic polyester as wetting agents. Additional wetting agent and/or oil will frequently be added to a tank mix for to facilitate dispersion on the foliage of the plant.

Other useful formulations for insecticidal applications are emulsifiable concentrates (ECs) which are homogeneous liquid compositions dispersible in water

or other dispersant, and may consist entirely of the insecticidal compound and a liquid or solid emulsifying agent, or may also contain a liquid carrier, such as xylene, heavy aromatic naphthas, isphorone, or other non-volatile organic solvents. For insecticidal application these concentrates are dispersed in water or other liquid carrier and normally applied as a spray to the area to be treated. The percentage by weight of the essential active ingredient may vary according to the manner in which the composition is to be applied, but in general comprises 0.5 to 95% of active ingredient by weight of the insecticidal composition.

Flowable formulations are similar to ECs, except that the active ingredient is suspended in a liquid carrier, generally water. Flowables, like ECs, may include a small amount of a surfactant, and will typically contain active ingredients in the range of 0.5 to 95%, frequently from 10 to 50%, by weight of the composition. For application, flowables may be diluted in water or other liquid vehicle, and are normally applied as a spray to the area to be treated.

Typical wetting, dispersing or emulsifying agents used in agricultural formulations include, but are not limited to, the alkyl and alkylaryl sulfonates and sulfates and their sodium salts; alkylaryl polyether alcohols; sulfated higher alcohols; polyethylene oxides; sulfonated animal and vegetable oils; sulfonated petroleum oils; fatty acid esters of polyhydric alcohols and the ethylene oxide addition products of such esters; and the addition product of long-chain mercaptans and ethylene oxide. Many other types of useful surface-active agents are available in commerce. Surface-active agents, when used, normally comprise 1 to 15% by weight of the composition.

Other useful formulations include suspensions of the active ingredient in a relatively non-volatile solvent such as water, corn oil, kerosene, propylene glycol, or other suitable solvents.

Still other useful formulations for insecticidal applications include simple solutions of the active ingredient in a solvent in which it is completely soluble at the desired concentration, such as acetone, alkylated naphthalenes, xylene, or other organic solvents. Granular formulations, wherein the toxicant is carried on relative coarse particles, are of particular utility for aerial distribution or for penetration of cover crop canopy. Pressurized sprays, typically aerosols wherein the active

ingredient is dispersed in finely divided form as a result of vaporization of a low-boiling dispersant solvent carrier may also be used. Water-soluble or water-dispersible granules are free flowing, non-dusty, and readily water-soluble or water-miscible. In use by the farmer on the field, the granular formulations, emulsifiable concentrates, flowable concentrates, aqueous emulsions, solutions, etc., may be diluted with water to give a concentration of active ingredient in the range of say 0.1% or 0.2% to 1.5% or 2%.

The active insecticidal compounds of this invention may be formulated and/or applied with one or more second compounds. Such combinations may provide certain advantages, such as, without limitation, exhibiting synergistic effects for greater control of insect pests, reducing rates of application of insecticide thereby minimizing any impact to the environment and to worker safety, controlling a broader spectrum of insect pests, safening of crop plants to phytotoxicity, and improving tolerance by non-pest species, such as mammals and fish.

Second compounds include, without limitation, other pesticides, plant growth regulators, fertilizers, soil conditioners, or other agricultural chemicals. In applying an active compound of this invention, whether formulated alone or with other agricultural chemicals, an effective amount and concentration of the active compound is of course employed; the amount may vary in the range of, e.g. about 0.001 to about 3 kg/ha, preferably about 0.03 to about 1 kg/ha. For field use, where there are losses of insecticide, higher application rates (e.g., four times the rates mentioned above) may be employed.

When the active insecticidal compounds of the present invention are used in combination with one or more of second compounds, e.g., with other pesticides such as herbicides, the herbicides include, without limitation, for example: N-(phosphonomethyl)glycine ("glyphosate"); aryloxyalkanoic acids such as (2,4-dichlorophenoxy)acetic acid ("2,4-D"), (4-chloro-2-methylphenoxy)acetic acid ("MCPA"), (+/-)-2-(4chloro-2-methylphenoxy)propanoic acid ("MCPP"); ureas such as N,N-dimethyl-N'-[4-(1-methylethyl)phenyl]urea ("isoproturon"); imidazolinones such as 2-[4,5-dihydro-4-methyl-4-(1-methylethyl)-5-oxo-1H-imidazol-2-yl]-3-pyridinecarboxylic acid ("imazapyr"), a reaction product comprising (+/-)-2-[4,5-dihydro-4-methyl-4-(1-methylethyl)-5-oxo-1H-imidazol-2-

yl]-4-methylbenzoic acid and (+/-)-2-[4,5-dihydro-4-methyl-4-(1-methylethyl)-5-oxo-1H-imidazol-2-yl]-5-methylbenzoic acid ("imazamethabenz"), (+/-)-2-[4,5-dihydro-4-methyl-4-(1-methylethyl)-5-oxo-1H-imidazol-2-yl]-5-ethyl-3-pyridinecarboxylic acid ("imazethapyr"), and (+/-)-2-[4,5-dihydro-4-methyl-4-(1-methylethyl)-5-oxo-1H-imidazol-2-yl]-3-quinolinecarboxylic acid ("imazaquin");

5 diphenyl ethers such as 5-[2-chloro-4-(trifluoromethyl)phenoxy]-2-nitrobenzoic acid ("acifluorfen"), methyl 5-(2,4-dichlorophenoxy)-2-nitrobenzoate ("bifenox"), and 5-[2-chloro-4-(trifluoromethyl)phenoxy]-N-(methylsulfonyl)-2-nitrobenzamide ("fomasafen"); hydroxybenzonitriles such as 4-hydroxy-3,5-diiodobenzonitrile

10 ("ioxynil") and 3,5-dibromo-4-hydroxybenzonitrile ("bromoxynil"); sulfonyleureas such as 2-[[[(4-chloro-6-methoxy-2-pyrimidinyl)amino]carbonyl]amino]sulfonyl]benzoic acid ("chlorimuron"), 2-chloro-N-[

15 [(4-methoxy-6-methyl-1,3,5-triazin-2-yl)amino]carbonyl]benzenesulfonamide (achlorsulfuron"), 2-[[[(4,6-dimethoxy-2-pyrimidinyl)amino]carbonyl]amino]sulfonyl]methyl]benzoic acid ("bensulfuron"), 2-[[[(4,6-dimethoxy-2-pyrimidinyl)amino]carbonyl]amino]sulfonyl]-1-methyl-1H-pyrazol-4-carboxylic acid ("pyrazosulfuron"), 3-[[[(4-methoxy-6-methyl-1,3,5-triazin-2-yl)amino]carbonyl]amino]sulfonyl]-2-thiophenecarboxylic acid

20 ("thifensulfuron"), and 2-(2-chloroethoxy)-N[[[(4-methoxy-6-methyl-1,3,5-triazin-2-yl)amino]carbonyl]benzenesulfonamide ("triasulfuron"); 2-(4-aryloxyphenoxy)alkanoic acids such as (+/-)-2[4-[(6-chloro-2-benzoxazolyl)oxy]phenoxy]propanoic acid ("fenoxaprop"), (+/-)-2-[4[[5-(trifluoromethyl)-2-pyridinyl]oxy]phenoxy]propanoic acid ("fluazifop"), (+/-)-2-[4-(6-chloro-2-quinoxalinyloxy]phenoxy]propanoic acid ("quizalofop"), and (+/-)-2-[(2,4-

25 dichlorophenoxy)phenoxy]propanoic acid ("diclofop"); benzothiadiazinones such as 3-(1-methylethyl)-1H-1,2,3-benzothiadiazin-4(3H)-one-2,2-dioxide ("bentazone"); 2-chloroacetanilides such as N-(butoxymethyl)-2-chloro-N-(2,6-diethylphenyl)acetamide ("butachlor"), 2-chloro-N-(2-ethyl-6-methylphenyl)-N-(2-methoxy-1-methylethyl)acetamide ("metolachlor"), 2-chloro-N-(ethoxymethyl)-N-

30 (2-ethyl-6-methylphenyl)acetamide ("acetochlor"), and (RS)-2-chloro-N-(2,4-dimethyl-3-thienyl)-N-(2-methoxy-1-methylethyl)acetamide ("dimethenamide"); arenecarboxylic acids such as 3,6-dichloro-2-methoxybenzoic acid ("dicamba");

pyridyloxyacetic acids such as [(4-amino-3,5-dichloro-6-fluoro-2-pyridinyl)oxy]acetic acid ("fluroxypyr"), and other herbicides.

When the active insecticidal compounds of the present invention are used in combination with one or more of second compounds, e.g., with other pesticides such as other insecticides, the other insecticides include, for example: organophosphate insecticides, such as chlorpyrifos, diazinon, dimethoate, malathion, parathion-methyl, and terbufos; pyrethroid insecticides, such as fenvalerate, deltamethrin, fenpropathrin, cyfluthrin, flucythrinate, *alpha*-cypermethrin, biphenthrin, resolved cyhalothrin, etofenprox, esfenvalerate, tralomehtrin, tefluthrin, cycloprothrin, betacyfluthrin, and acrinathrin; carbamate insecticides, such as aldecarb, carbaryl, carbofuran, and methomyl; organochlorine insecticides, such as endosulfan, endrin, heptachlor, and lindane; benzoylurea insecticides, such as diflubenuron, triflumuron, teflubenzuron, chlorfluazuron, flucycloxuron, hexaflumuron, flufenoxuron, and lufenuron; and other insecticides, such as amitraz, clofentezine, fenpyroximate, hexythiazox, spinosad, and imidacloprid.

When the active insecticidal compounds of the present invention are used in combination with one or more of second compounds, e.g., with other pesticides such as fungicides, the fungicides include, for example: benzimidazole fungicides, such as benomyl, carbendazim, thiabendazole, and thiophanate-methyl; 1,2,4-triazole fungicides, such as epoxyconazole, cyproconazole, flusilazole, flutriafol, propiconazole, tebuconazole, triadimefon, and triadimenol; substituted anilide fungicides, such as metalaxyl, oxadixyl, procymidone, and vinclozolin; organophosphorus fungicides, such as fosetyl, iprobenfos, pyrazophos, edifenphos, and tolclofos-methyl; morpholine fungicides, such as fenpropimorph, tridemorph, and dodemorph; other systemic fungicides, such as fenarimol, imazalil, prochloraz, tricyclazole, and triforine; dithiocarbamate fungicides, such as mancozeb, maneb, propineb, zineb, and ziram; non-systemic fungicides, such as chlorothalonil, dichlofluanid, dithianon, and iprodione, captan, dinocap, dodine, fluazinam, gluazatine, PCNB, pencycuron, quintozone, tricylamide, and validamycin; inorganic fungicides, such as copper and sulphur products, and other fungicides.

When the active insecticidal compounds of the present invention are used in combination with one or more of second compounds, e.g., with other pesticides such

as nematicides, the nematicides include, for example: carbofuran, carbosulfan, turbufos, aldecarb, ethoprop, fenamphos, oxamyl, isazofos, cadusafos, and other nematicides.

When the active insecticidal compounds of the present invention are used in combination with one or more of second compounds, e.g., with other materials such as plant growth regulators, the plant growth regulators include, for example: maleic hydrazide, chlormequat, ethephon, gibberellin, mepiquat, thidiazon, inabenfide, triaphenthenol, paclobutrazol, uniconazole, DCPA, prohexadione, trinexapac-ethyl, and other plant growth regulators.

Soil conditioners are materials which, when added to the soil, promote a variety of benefits for the efficacious growth of plants. Soil conditioners are used to reduce soil compaction, promote and increase effectiveness of drainage, improve soil permeability, promote optimum plant nutrient content in the soil, and promote better pesticide and fertilizer incorporation. When the active insecticidal compounds of the present invention are used in combination with one or more of second compounds, e.g., with other materials such as soil conditioners, the soil conditioners include organic matter, such as humus, which promotes retention of cation plant nutrients in the soil; mixtures of cation nutrients, such as calcium, magnesium, potash, sodium, and hydrogen complexes; or microorganism compositions which promote conditions in the soil favorable to plant growth. Such microorganism compositions include, for example, *bacillus*, *pseudomonas*, *azotobacter*, *azospirillum*, *rhizobium*, and soil-borne *cyanobacteria*.

Fertilizers are plant food supplements, which commonly contain nitrogen, phosphorus, and potassium. When the active insecticidal compounds of the present invention are used in combination with one or more of second compounds, e.g., with other materials such as fertilizers, the fertilizers include nitrogen fertilizers, such as ammonium sulfate, ammonium nitrate, and bone meal; phosphate fertilizers, such as superphosphate, triple superphosphate, ammonium sulfate, and diammonium sulfate; and potassium fertilizers, such as muriate of potash, potassium sulfate, and potassium nitrate, and other fertilizers.

The following examples further illustrate the present invention, but, of course, should not be construed as in any way limiting its scope. The examples are

organized to present protocols for the synthesis of the compounds of formula I of the present invention, set forth a list of such synthesized species, and set forth certain biological data indicating the efficacy of such compounds.

Example 1

- 5 This example illustrates the preparation of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-4-hydroxy-1-[(4-pyrimidin-2-yloxyphenyl)methyl]piperidin-1-oxide (Compound 112 in table below)

Step A Synthesis of 2-[4-(chloromethyl)phenoxy]pyrimidine as an

10 Intermediate

- A stirred solution of 2.0 grams (0.0099 mole) of (4-pyrimidin-2-yloxyphenyl)methan-1-ol (known compound) and 7 drops of pyridine in 50 mL of methylene chloride was cooled in an ice-water bath, and 0.94 mL (0.013 mole) of thionyl chloride was added dropwise. Upon completion of the addition, the reaction mixture was stirred for 3 hours at 10°C to 20°C. The reaction mixture was then poured into ice-water and basified using sodium bicarbonate. The aqueous layer was separated from the organic layer, and was extracted one time with 75 mL of methylene chloride. The methylene chloride extract and organic layer were combined and passed through silicone coated filter paper. The filtrate was then concentrated under reduced pressure, yielding 2.1 grams of the subject compound.

Step B Synthesis of 1-[(4-pyrimidin-2-yloxyphenyl)methyl]piperidin-4-one as an Intermediate

- A solution of 1.47 grams (0.0096 mole) of 4-piperidone monohydrate hydrochloride, 2.1 grams (0.0096 mole) of 2-[4-(chloromethyl)phenoxy]pyrimidine, and 4.34 grams (0.0336 mole) of bis(methylethyl)ethylamine in 35 mL of dimethylsulfoxide (DMSO) was stirred at ambient temperature for about 24 hours. The reaction mixture was then diluted with 200 mL of water and was extracted two times with 200 mL of ethyl acetate. The extracts were then combined and washed two times with 75 mL of an aqueous mixture of 10% lithium chloride. The resultant organic layer was dried over magnesium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was purified with column chromatography on silica gel using 1:2 ethyl acetate:petroleum ether as an eluant.

The appropriate fractions were combined and concentrated under reduced pressure, yielding 2.71 grams of the subject compound.

Step C Synthesis of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-1-[(4-pyrimidin-2-yloxyphenyl)methyl]piperidin-4-ol as an Intermediate

5 A stirred solution of 0.8 gram (0.0019 mole) of bis(trifluoromethoxyphenyl)bromomethane and 2.2 grams (0.0078 mole) of 1-[(4-pyrimidin-2-yloxyphenyl)methyl]piperidin-4-one in 50 mL of THF was chilled to -78°C, and 1.5 mL of n-butyl lithium (2.5 M) was added dropwise during a 15-minute period while maintaining the temperature of the reaction mixture between -80°C to -70°C. The reaction mixture was then allowed to warm to about 0°C. The reaction mixture was quenched with a saturated aqueous solution of ammonium chloride, and the mixture was extracted two times with 75 mL of ethyl acetate. The combined extracts were dried over sodium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was purified with column chromatography on silica gel using 1:4 acetone:methylene chloride as an eluant. The appropriate fractions were combined and further purified by passing them through a Waters SEP-PAK® Vac 35cc NH₂ Cartridge (purchased from Waters, 34 Maple Street, Milford, MA 01757) using 1:4 ethyl acetate:petroleum ether as an eluant. The appropriate fractions were combined and concentrated under reduced pressure, yielding 0.42 gram of the subject compound. The NMR spectrum was consistent with the proposed structure.

25 Step D Synthesis of Compound 112

 A solution of 0.28 gram (0.00045 mole) of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-1-[(4-pyrimidin-2-yloxyphenyl)methyl]piperidin-4-ol and 1.4 grams of 50% aqueous hydrogen peroxide in 40 mL of methanol was stirred at ambient temperature for about 4 days. The reaction mixture was then diluted with 200 mL of water and extracted twice with 200 mL each of ethyl acetate. The combined extracts were then washed twice with 75 mL each of aqueous mixture of 10% lithium chloride. The organic layer was dried over magnesium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was

purified with column chromatography on silica gel using 1:2 ethyl acetate:petroleum ether as an eluant. The appropriate fractions were combined and concentrated under reduced pressure, yielding 0.29 gram of Compound 112.

5

Example 2

This example illustrates the preparation of 4-{bis[4-(trifluoromethoxy)phenyl]hydroxymethyl}-1-[[4-(2-methyl(1,2,3,4-tetrazol-5-yl))phenyl]methyl]piperidin-4-ol (Compound 93 in the table below)

- 10 Step A Synthesis of 5-[4-(bromomethyl)phenyl]-2-methyl-1,2,3,4-tetrazole
 as an Intermediate
 A stirred solution of 45 grams (0.258 mole) of 2-methyl-5-(4-methylphenyl)-
 1,2,3,4-tetrazole (known compound), 46 grams (1 equivalent) of N-
 bromosuccinimide, and a catalytic amount of benzoyl peroxide in 200 mL of carbon
15 tetrachloride was irradiated with light during a 3.5 hour period. The mixture was
 then cooled in an ice bath and filtered to collect 35.1 grams of the subject
 compound. The NMR spectrum was consistent with the proposed structure.
- 20 Step B Synthesis of 1-[[4-(2-methyl-1,2,3,4-tetrazol-5-
 yl)phenyl]methyl]piperidin-4-one as an Intermediate
 A solution of 23 grams (0.091mole) of 5-[4-(bromomethyl)phenyl]-2-
 methyl-1,2,3,4-tetrazole, 14 grams (0.091 mole) of 4-piperidone monohydrate
 hydrochloride, and 47 mL (3 equivalents) of N,N-diisopropylethylamine in 200 mL
25 of dimethylsulfoxide (DMSO) was stirred for about 3 days. The reaction was
 quenched by pouring the reaction mixture onto 400 mL of dilute, cold sodium
 hydroxide. The resultant solution was extracted one time with 300 mL of ethyl
 acetate. An emulsion formed, which was broken up by warming it to about 35°C.
 The organic layer was separated and concentrated under reduced pressure to a
 residue. The residue was purified with column chromatography on silica gel using
30 40:1 methylene chloride:methanol as an eluant. The appropriate fractions were
 combined and concentrated under reduced pressure, yielding 9.12 grams of the
 subject compound. The NMR spectrum was consistent with the proposed structure.
- 35 Step C Synthesis of 4-hydroxy-1-[[4-(2-methyl(1,2,3,4-tetrazol-5-
 yl))phenyl]methyl]piperidine-4-carbonitrile as an Intermediate

A stirred solution of 8.62 grams (0.032 mole) of 1-{{[4-(2-methyl-1,2,3,4-tetrazol-5-yl))phenyl]methyl}piperidin-4-one in 100 mL of ether and 150 mL of water was cooled to about 10°C and 3.9 grams (2.5 equivalents) of sodium cyanide was added in one portion. To this was added 6.6 mL (2.5 equivalents) of hydrochloric acid (12 M) dropwise while maintaining the reaction mixture temperature at about 10°C. Upon completion of the addition, the reaction mixture was stirred for 1.5 hours while warming to ambient temperature. The reaction mixture was then poured into 200 mL of water, to which was then added 200 mL of ethyl acetate. The organic layer was separated and concentrated under reduced pressure to yield 9.6 grams of the subject compound. The NMR spectrum was consistent with the proposed structure.

Step D Synthesis of ethyl 4-hydroxy-1-{{[4-(2-methyl(1,2,3,4-tetrazol-5-yl))phenyl]methyl}piperidine-4-carboxylate as an Intermediate

A solution of 9.6 grams (0.032 mole) of 4-hydroxy-1-{{[4-(2-methyl(1,2,3,4-tetrazol-5-yl))phenyl]methyl}piperidine-4-carbonitrile in 200 mL of ethanol was saturated with gaseous hydrogen chloride, and then it was stirred at 55°C for about 20 hours. The reaction mixture was allowed to cool and then it was poured onto 500 mL of ice. The resultant mixture was basified with 50% sodium hydroxide and extracted once with 300 mL of ethyl acetate. The extract was washed three times with 80 mL brine, dried over magnesium sulfate, filtered and concentrated under reduced pressure to a residue, yielding 5.14 grams of the subject compound. The NMR spectrum was consistent with the proposed structure.

Step E Synthesis of Compound 93

A solution of 1.6 grams (4.5 equivalents) of magnesium and 8.9 mL (4 equivalents) of 4-trifluoromethoxybromobenzene (known compound) in 75 mL of THF was stirred and 5.14 grams (0.0149 mole) of ethyl 4-hydroxy-1-{{[4-(2-methyl(1,2,3,4-tetrazol-5-yl))phenyl]methyl}piperidine-4-carboxylate was added in one portion. After a mild exotherm, the reaction mixture was heated to 50°C where it was stirred for 2 hours. The reaction mixture was allowed to cool to ambient temperature as it stirred for 72 hours; then it was poured into 200 mL of an aqueous solution saturated with ammonium chloride. The mixture was extracted one time

with 200 mL of ethyl acetate, and the extract was washed two times with 80 mL of brine. The extract was dried with magnesium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was purified with column chromatography on silica gel using diethyl ether as an eluant. The appropriate
5 fractions were combined and concentrated under reduced pressure, yielding 9.29 grams of Compound 93.

Example 3

This example illustrates the preparation of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-4-fluoro-1-[(4-pyrimidin-2-yloxyphenyl)methyl]piperidine (Compound 192 in the table below)
10

A stirred solution of 0.11 gram (0.170 mmole) of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-1-[(4-pyrimidin-2-yloxyphenyl)methyl]piperidin-
15 4-ol (prepared as in Example 1, Steps A through C) in 2.0 mL of methylene chloride was cooled to -40°C, and 18.2 µL (0.186 mmoles) of (dimethylamino)sulfur trifluoride was added. The reaction mixture was allowed to warm to ambient temperature where it stirred for 20 minutes. The reaction mixture was then poured onto 10 mL of an aqueous solution saturated with sodium bicarbonate, and the
20 mixture was extracted with three 20 mL portions of ethyl acetate. The combined extracts were dried with sodium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was purified with column chromatography on silica gel using 1:1 ethyl acetate:hexane as an eluant. The appropriate fractions were combined and concentrated under reduced pressure, yielding 85 milligrams of
25 Compound 192. The NMR spectrum was consistent with the proposed structure.

30

Example 4

This example illustrates the preparation of ethyl 4-{bis[4-(trifluoromethyl)phenyl]chloromethyl}-1-[(4-(2-pyridyloxy)phenyl)methyl]piperidine-4-carboxylate (Compound 195 in the table below)
35

Step A Synthesis of ethyl 1-[(tert-butyl)oxycarbonyl]-4-{bis[4-(trifluoromethyl)phenyl]hydroxymethyl}piperidine-4-carboxylate as an Intermediate

A stirred solution of 1 mL of diisopropyl amine in 10 mL THF was cooled to about 0°C and 2.55 mL of n-butyl lithium (2.5 M in hexane) was added slowly. The reaction mixture was stirred for 15 minutes, then it was cooled to -78°C. To this was then added a solution of 1.26 grams (4.9 mmole) of ethyl 1-[(tert-butyl)oxycarbonyl]piperidine-4-carboxylate (known compound) in 10 mL of THF. The reaction mixture continued to stir at -78°C for 1 hour, then a solution of 1.56 grams (4.9 mmole) of di-4-(trifluoromethyl)phenyl ketone (known compound) in 5 mL of THF was added. Upon completion of the addition, the reaction mixture was warmed to ambient temperature during a 14 hour period. The reaction was then quenched by adding 125 mL of aqueous 5% hydrochloric acid to the reaction mixture. The mixture was then extracted with three 125 mL portions of ethyl acetate and the combined extracts were dried with sodium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was purified with column chromatography on silica gel using 1:5 ethyl acetate:hexane as an eluant. The appropriate fractions were combined and concentrated under reduced pressure, yielding 2.2 grams of the subject compound. The NMR spectrum was consistent with the proposed structure.

Step B Synthesis of ethyl 1-[(tert-butyl)oxycarbonyl]-4-{bis[4-(trifluoromethyl)phenyl]chloromethyl}piperidine-4-carboxylate as an Intermediate

A solution of 290 milligrams (0.504 mmole) of ethyl 1-[(tert-butyl)oxycarbonyl]-4-{bis[4-(trifluoromethyl)phenyl]hydroxymethyl}piperidine-4-carboxylate, 110 µL (1.513 mmole) of thionyl chloride, and 408 µL (5.04 mmole) of pyridine in 5 mL of methylene chloride was stirred at ambient temperature for 1 hour. The reaction mixture was then poured into 25 mL of an aqueous solution saturated with sodium bicarbonate, and the mixture was extracted with three 50 mL portions of ethyl acetate. The combined extracts were dried with sodium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was purified with column chromatography on silica gel using 1:10 ethyl acetate:hexane as an eluant. The appropriate fractions were combined and concentrated under

reduced pressure, yielding 240 milligrams of the subject compound. The NMR spectrum was consistent with the proposed structure.

5 Step C Synthesis of ethyl 4-{bis[4-(trifluoromethyl)phenyl]chloromethyl}piperidine-4-carboxylate as an Intermediate

A solution of 190 milligrams (0.320 mmole) of ethyl 1-[(tert-butyl)oxycarbonyl]-4-{bis[4-(trifluoromethyl)phenyl]chloromethyl}piperidine-4-carboxylate and 2 mL of trifluoro acetic acid (TFA) in 0.5 mL of methylene chloride
10 was stirred at room temperature for 10 minutes. The reaction mixture was then diluted with toluene and concentrated under reduced pressure, yielding 158 milligrams of the subject compound. The NMR spectrum was consistent with the proposed structure.

15 Step D Synthesis of Compound 195

A solution of 158 milligrams (0.320 mmole) of ethyl 4-{bis[4-(trifluoromethyl)phenyl]chloromethyl}piperidine-4-carboxylate, 78 milligrams (0.352 mmole) of 2-[4-(chloromethyl)phenoxy]pyridine and 0.28 mL of diisopropylethylamine in 4 mL of DMF was stirred at room temperature for 48
20 hours. The reaction mixture was then poured into 20 mL of an aqueous solution saturated with sodium bicarbonate, and the mixture was extracted with three 40 mL portions of ethyl acetate. The combined extracts were dried with sodium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was purified with column chromatography on silica gel using 1:5 ethyl acetate:hexane as
25 an eluant. The appropriate fractions were combined and concentrated under reduced pressure, yielding 42 milligrams of Compound 195. The NMR spectrum was consistent with the proposed structure.

30 Example 5

This example illustrates the preparation of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-4-hydroxy-1-{[4-(2-methoxyethoxy)phenyl]methyl}piperidin-1-oxide (Compound 385 in table below)

35 Step A Synthesis of 4-(2-methoxyethoxy)benzaldehyde as an Intermediate

A solution of 5.0 grams (0.041 mole) of 4-hydroxybenzaldehyde, 9.5 grams (0.041 mole) of *p*-toluene sulfonic acid 2-methoxyethyl ester, and 6.3 grams (0.046 mole) of anhydrous potassium carbonate in 50 mL of dimethylsulfoxide (DMSO) was stirred at ambient temperature for about 72 hours. The reaction mixture was then partitioned between 300 mL of an aqueous mixture of 10% lithium chloride and 100 mL of ethyl acetate. The ethyl acetate portion was washed three times; first, with 100 mL of an aqueous mixture of 10% sodium hydroxide, second, with 100 mL of an aqueous mixture of 10% lithium chloride, and, third, with 100 mL of brine. The resultant organic layer was dried over magnesium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was purified with column chromatography on silica gel using methylene chloride as an eluant. The appropriate fractions were combined and concentrated under reduced pressure, yielding 7.3 grams of the subject compound. The NMR spectrum was consistent with the proposed structure.

Step B Synthesis of tert-butyl 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-4-hydroxypiperidine carboxylate as an Intermediate

A stirred solution of 8.9 grams (0.022 mole) of bis(trifluoromethoxyphenyl)bromomethane and 4.82 grams (0.024 mole) of tert-butyl 4-oxo-1-piperidine carboxylate in 150 mL of THF was chilled to -78°C, and 9.24 mL of n-butyl lithium (2.5 M) was added dropwise during a 15-minute period while maintaining the temperature of the reaction mixture between -85°C to -75°C. The reaction mixture was then stirred for 30 minutes while maintaining the temperature of the reaction mixture between -80°C to -70°C. The reaction mixture was quenched with an aqueous solution saturated with ammonium chloride keeping the internal temperature at less than -55°C, then the mixture was extracted two times with 100 mL of ethyl acetate. The extracts were combined and washed once with 75 mL of an aqueous mixture of 10% lithium chloride. The combined extracts were dried over sodium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was triturated with petroleum ether and a trace of ether then filtered, yielding 3.43 grams of the subject compound. The NMR spectrum was consistent with the proposed structure.

Step C Synthesis of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}piperidin-4-ol as an Intermediate

5 A mixture of 3.0 grams (0.0057 mole) of tert-butyl 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-4-hydroxypiperidine carboxylate and 25 mL of methylene chloride was chilled in a wet ice bath. A solution of 2.63 mL trifluoroacetic acid and 5 mL methylene chloride was added dropwise to the reaction mixture during a 5-minute period. The wet ice bath was removed and the reaction mixture was allowed to warm to ambient temperature. The reaction mixture was then warmed to reflux, and allowed to cool to ambient temperature. The reaction mixture was concentrated under reduced pressure. The residue can then be dissolved in 100 mL methylene chloride and reacted with 200 mL of an aqueous solution saturated with sodium carbonate. The methylene chloride layer can then be separated, dried over magnesium sulfate, filtered and concentrated under reduced pressure to yield the subject compound.

Step D Synthesis of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-1-{[4-(2-methoxyethoxy)phenyl]methyl}piperidin-4-ol (Compound 308 in table below) as an Intermediate

20 A solution of 2.2 grams (0.0050 mole) of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}piperidin-4-ol, 1.1 grams (0.0061 mole) of 4-(2-methoxyethoxy)benzaldehyde and 1.35 grams (0.0064 mole) of sodium triacetoxyborohydride in 25 mL of methylene chloride was stirred at ambient temperature for 18 hours. The reaction mixture was then diluted with 200 mL of water and stirred at ambient temperature for 5 hours. The phases were separated. The organic phase was washed three times; first, with 100 mL of an aqueous mixture of 10% sodium hydroxide, second, with 100 mL of an aqueous mixture of 10% lithium chloride, and, third, with 100 mL of brine. The resultant organic layer was dried over magnesium sulfate, filtered and concentrated under reduced pressure to a residue. The residue was purified with column chromatography on silica gel using methylene chloride:1%-5% methanol as an eluant. The appropriate fractions were combined and concentrated under reduced pressure, yielding 2.2 grams of the subject compound. The NMR spectrum was consistent with the proposed structure.

Step E Synthesis of Compound 385

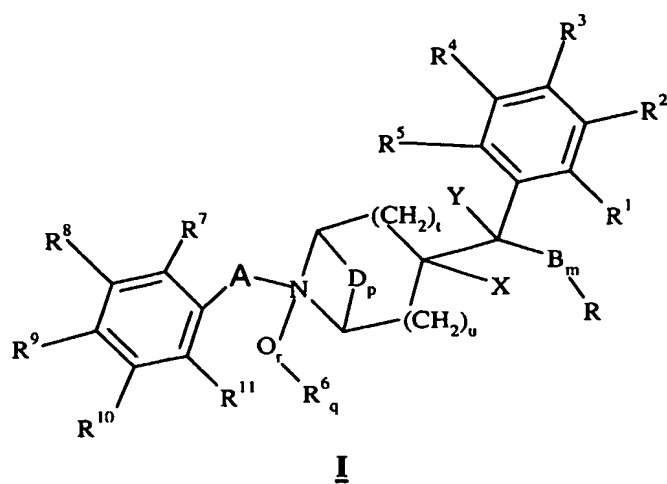
5 A solution of 1.9 grams (0.0032 mole) of 4-{bis[4-(trifluoromethoxy)phenyl]methyl}-1-[[4-(2-methoxyethoxy)phenyl]methyl]piperidin-4-ol and 2.0 mL of 50% aqueous hydrogen peroxide in 25 mL of methanol was stirred at ambient temperature for about 7 days. The reaction mixture was then partitioned between 300 mL of an aqueous mixture of 10% lithium chloride and 100 mL of ethyl acetate. The aqueous phase was extracted twice with 100 mL each of ethyl acetate. The organic phase and extracts were combined and washed twice; first, with 100 mL of aqueous mixture of 10% lithium chloride, second, with 100 mL of brine. The resultant organic layer was dried over sodium sulfate, filtered and concentrated under reduced pressure to a residue, yielding 1.9 gram of Compound 385. The NMR spectrum was consistent with the proposed structure.

15 It is well known to one of ordinary skill in the art that compounds like the compounds of formula I of the present invention can contain optically active and racemic forms. It is also well known in the art that compounds like the compounds of formula I may contain stereoisomeric forms, tautomeric forms and/or exhibit polymorphism. It is to be understood that the present invention encompasses any racemic, optically active, polymorphic, tautomeric, or stereoisomeric form, or mixtures thereof. It should be noted that it is well known in the art how to prepare optically active forms, for example by resolution of a racemic mixture, or by synthesis from optically active intermediates.

20 The following table sets forth some additional examples of compounds of formula I.

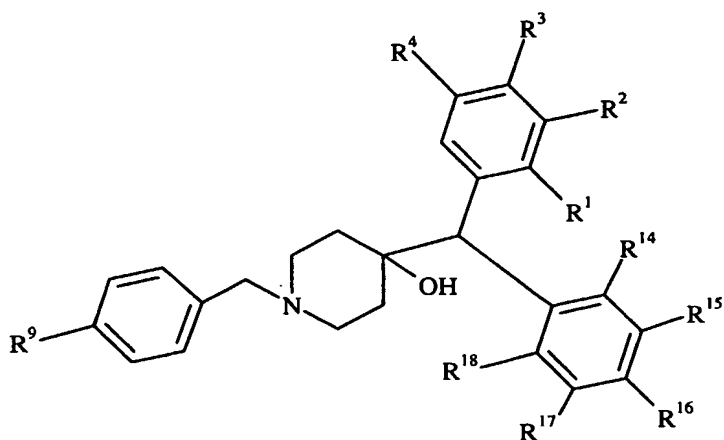
Table 1
N-Substituted Azacycles

58



Where

- 5 m, p, q and r are 0; t and u are 1; R is phenyl substituted with R^{14} , R^{15} , R^{16} , R^{17} , R^{18} ; X is OH; A is $-\text{CH}_2-$; and R^5 , R^7 , R^8 , R^{10} , R^{11} and Y are H:



I - A

Compd No.	R ¹	R ²	R ³	R ⁴	R ⁹	R ¹⁴	R ¹⁵	R ¹⁶	R ¹⁷	R ¹⁸
1	Cl	H	H	H	pyrimidin-2-yloxy	H	H	H	H	Cl
2	H	H	CF ₃	H	pyrimidin-2-yloxy	H	H	OCF ₃	H	H
3	H	Cl	H	Cl	pyrimidin-2-yloxy	H	H	CF ₃	H	H
4	H	F	Cl	H	pyrimidin-2-yloxy	H	H	Cl	F	H
5	H	H	CF ₃	H	pyrimidin-2-yloxy	H	OCH ₃	H	OCH ₃	H
6	H	Cl	Cl	H	pyrimidin-2-yloxy	H	Cl	Cl	H	H
7	H	H	CF ₃	H	pyrimidin-2-yloxy	H	Cl	Cl	H	H
8	H	F	H	H	pyrimidin-2-yloxy	H	H	H	F	H
9	H	Cl	Cl	H	pyrimidin-2-yloxy	H	H	CF ₃	F	H
10	H	F	H	F	pyrimidin-2-yloxy	H	F	H	H	H
11	H	F	H	F	pyrimidin-2-yloxy	H	H	CF ₃	F	H
12	H	H	CF ₃	H	pyrimidin-2-yloxy	H	H	H	H	H
13	H	H	CF ₃	H	pyrimidin-2-yloxy	H	H	H	Cl	H
14	H	H	CF ₃	H	pyrimidin-2-yloxy	Cl	H	H	Cl	H
15	Cl	H	Cl	H	pyrimidin-2-yloxy	H	H	CF ₃	H	H
16	H	H	CF ₃	H	pyrimidin-2-yloxy	H	H	CF ₃	H	H
17	H	H	CF ₃	H	pyrimidin-2-yloxy	H	H	CF ₃	F	H
18	H	H	CF ₃	H	pyrimidin-2-yloxy	H	H	H	H	H
19	H	H	OCF ₃	H	pyrimidin-2-yloxy	H	H	OCF ₃	H	Cl
20	H	H	OCF ₃	H	pyrimidin-2-yl	H	H	OCF ₃	H	H
21	H	H	OCF ₃	H	pyrimidin-2-yloxy	H	H	OCF ₃	H	H
22	H	H	CF ₃	H	pyrid-2-yloxy	H	H	Cl	H	H
23	H	H	Cl	H	pyrid-2-yloxy	H	H	H	H	H
24	H	H	H	Cl	2-ethyl-2H-tetrazol-5-yl	H	H	H	Cl	H
25	H	H	OCF ₃	H	2-ethyl-2H-tetrazol-5-yl	H	H	OCF ₃	H	H
26	H	Cl	Cl	H	2-ethyl-2H-tetrazol-5-yl	H	H	Cl	Cl	H
27	H	H	CF ₃	H	2-ethyl-2H-tetrazol-5-yl	H	H	CF ₃	H	H
28	H	H	OCF ₃	H	2-ethyl-2H-tetrazol-5-yl	H	H	OCF ₃	H	H
29	H	H	OCF ₃	H	CH=NOC ₂ H ₅ NHC(=O)OCH(CH ₃) ₂	H	H	OCF ₃	H	H

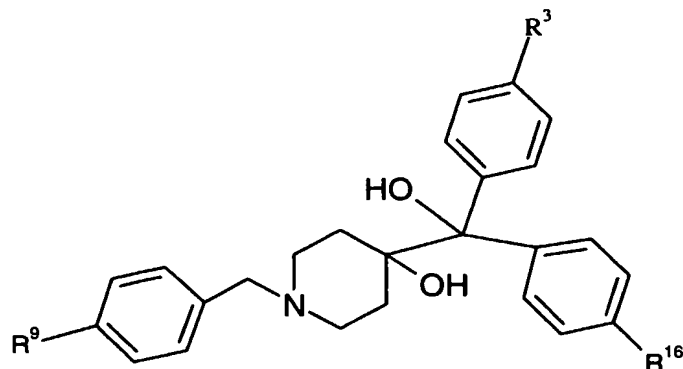
Compd No.	R ¹	R ²	R ³	R ⁴	R ⁹	R ¹⁴	R ¹⁵	R ¹⁶	R ¹⁷	R ¹⁸
30	H	H	OCF ₃	H	3-(CF ₃)pyrid-2-ylloxy	H	H	OCF	H	H
31	H	H	CF ₃	H	CH=NOCH ₃	H	H	CF ₃	H	H
32	H	H	CF ₃	H	CH=NOCH ₃ H ₇	H	H	CF ₃	H	H
33	H	H	CF ₃	H	CH=NOCH(CH ₃) ₂	H	H	CF ₃	H	H
34	H	H	CF ₃	H	CH=NOCH ₂ CH=CH ₂	H	H	CF ₃	H	H
35	H	H	CF ₃	H	CH=NOCH ₂ C≡CH	H	H	CF ₃	H	H
36	H	H	CF ₃	H	CH=NOCH ₂ H ₅	H	H	CF ₃	H	H
37	H	H	OCF ₃	H	CH=NOCH ₃	H	H	OCF ₃	H	H
38	H	H	OCF ₃	H	CH=NOCH ₃ H ₇	H	H	OCF ₃	H	H
39	H	H	OCF ₃	H	CH=NOCH(CH ₃) ₂	H	H	OCF ₃	H	H
40	H	H	OCF ₃	H	CH=NOCH ₂ CH=CH ₂	H	H	OCF ₃	H	H
41	H	H	OCF ₃	H	CH=NOCH ₂ C≡CH	H	H	OCF ₃	H	H
42	H	H	OCF ₃	H	CH=NOCH ₂ H ₅	H	H	OCF ₃	H	H
43	H	H	CF ₃	H	CH=NOCH ₂ H ₅	F	H	CF ₃	H	H
44	H	H	CF ₃	H	CH=NOCH ₂ H ₅	Cl	H	CF ₃	H	H
45	H	H	OCF ₃	H	phenoxy	H	H	OCF ₃	H	H
46	H	H	OCF ₃	H	4-(1-methylethoxy)carbonyl)phenoxy	H	H	OCF ₃	H	H
47	H	H	OCF ₃	H	4-(methoxycarbonyl)phenoxy	H	H	OCF ₃	H	H
48	H	H	OCF ₃	H	6-chloro-pyridazin-3-ylloxy	H	H	OCF ₃	H	H
49	H	H	CF ₃	H	6-chloro-pyridazin-3-ylloxy	H	H	CF ₃	H	H
50	H	H	OCF ₃	H	NHCO ₂ CH ₃	H	H	OCF ₃	H	H
51	H	H	OCF ₃	H	NHCO ₂ C ₂ H ₅	H	H	OCF ₃	H	H
52	H	H	OCF ₃	H	NHCO ₂ CH ₂ CH=CH ₂	H	H	OCF ₃	H	H
53	H	H	OCF ₃	H	NHCO ₂ CH ₂ C≡CH	H	H	OCF ₃	H	H
54	H	H	CF ₃	H	NHCO ₂ CH(CH ₃) ₂	H	H	CF ₃	H	H
55	H	H	CF ₃	H	NHCO ₂ CH ₃	H	H	CF ₃	H	H
56	H	H	CF ₃	H	NHCO ₂ CH ₂ CH=CH ₂	H	H	CF ₃	H	H
57	H	H	CF ₃	H	OC(=O)NHCH ₃	H	H	CF ₃	H	H
58	H	H	CF ₃	H	OC(=O)NHC ₂ H ₅	H	H	CF ₃	H	H
59	H	H	CF ₃	H	OC(=O)NHCH(CH ₃) ₂	H	H	CF ₃	H	H
60	H	H	CF ₃	H	OC(=O)NHCH ₂ CH=CH ₂	H	H	CF ₃	H	H
61	H	H	OCF ₃	H	OC(=O)NHCH ₃	H	H	OCF ₃	H	H
62	H	H	OCF ₃	H	OC(=O)NHC ₂ H ₅	H	H	OCF ₃	H	H

Compd No.	R ¹	R ²	R ³	R ⁴	R ⁹	R ¹⁴	R ¹⁵	R ¹⁶	R ¹⁷	R ¹⁸
63	H	H	OCF ₃	H	OC(=O)NHCH(CH ₃) ₂	H	H	OCF ₃	H	H
64	H	H	OCF ₃	H	OC(=O)NHCH ₂ CH=CH ₂	H	H	OCF ₃	H	H
65	H	H	OCF ₃	H	OC(=O)NHCH ₂ C≡CH	H	H	OCF ₃	H	H
66	H	H	CF ₃	H	OCH ₃	H	H	CF ₃	H	H
67	H	H	CF ₃	H	OC ₂ H ₅	H	H	CF ₃	H	H
68	H	H	CF ₃	H	OC ₃ H ₇	H	H	CF ₃	H	H
69	H	H	CF ₃	H	cyclopentoxo	H	H	CF ₃	H	H
70	H	H	OCF ₃	H	OCH ₃	H	H	OCF ₃	H	H
71	H	H	OCF ₃	H	OC ₂ H ₅	H	H	OCF ₃	H	H
72	H	H	OCF ₃	H	OC ₃ H ₇	H	H	OCF ₃	H	H
73	H	H	OCF ₃	H	OCH(CH ₃) ₂	H	H	OCF ₃	H	H
74	H	H	OCF ₃	H	cyclopentoxo	H	H	OCF ₃	H	H
75	H	H	OCF ₂	H	pyrid-2-yloxy	H	H	OCF ₂	H	H
76	H	H	OCF ₂	H	pyrimidin-2-yloxy	H	H	OCF ₂	H	H
77	H	H	OCF ₂	H	CH=NOC ₂ H ₅	H	H	OCF ₂	H	H
78	H	H	OCF ₂	H	OC(=O)NHCH(CH ₃) ₂	H	H	OCF ₂	H	H
79	H	H	OCF ₂	H	NHCO ₂ CH(CH ₃) ₂	H	H	OCF ₂	H	H
80	H	H	OCF ₂	H	OC ₃ H ₇	H	H	OCF ₂	H	H
81	H	H	OCF ₂ CHF ₂	H	pyrid-2-yloxy	H	H	OCF ₂ CHF ₂	H	H
82	H	H	OCF ₂ CHF ₂	H	pyrimidin-2-yloxy	H	H	OCF ₂ CHF ₂	H	H
83	H	H	OCF ₂ CHF ₂	H	6-chloro-pyridazin-3-yloxy	H	H	OCF ₂ CHF ₂	H	H
84	H	H	OCF ₂ CHF ₂	H	OC ₃ H ₇	H	H	OCF ₂ CHF ₂	H	H
85	H	H	OCF ₂ CHF ₂	H	OC(=O)NHCH(CH ₃) ₂	H	H	OCF ₂ CHF ₂	H	H
86	H	H	OCF ₂ CHF ₂	H	NHCO ₂ CH(CH ₃) ₂	H	H	OCF ₂ CHF ₂	H	H
87	H	H	OCF ₂ CHF ₂	H	OCO ₂ CH(CH ₃) ₂	H	H	OCF ₂ CHF ₂	H	H
88	H	H	SF ₅	H	pyrimidin-2-yloxy	H	H	SF ₅	H	H
89	H	H	SF ₅	H	pyrimidin-2-yloxy	H	H	CF ₃	H	H
90	H	H	OCF ₃	H	CO ₂ CH(CH ₃) ₂	H	H	OCF ₃	H	H

where

m, p, q and r are 0; t and u are 1; R is phenyl substituted with R^{14} , R^{15} , R^{16} , R^{17} , R^{18} ; X and Y are OH; A is $-\text{CH}_2-$; and R^1 , R^2 , R^4 , R^5 , R^{14} , R^{15} , R^{17} , R^{18} , R^7 , R^8 , R^{10} , and R^{11} are H:

5



I - B

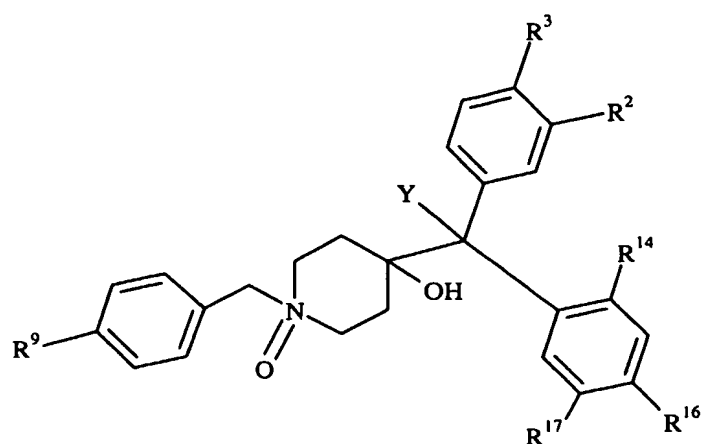
Compd. No.	R^3	R^9	R^{16}
91	OCF_3	OC_3H_7	OCF_3
92	CF_3	OC_3H_7	CF_3
93	OCF_3	2-methyl-2H-tetrazol-5-yl	OCF_3
94	CF_3	pyrid-2-yloxy	CF_3
95	CF_3	pyrimidin-2-yloxy	CF_3
96	CF_3	6-chloro-pyridazin-3yloxy	CF_3
97	CF_3	6-methoxy-pyridazin-3yloxy	CF_3
98	CF_3	2-ethyl-2H-tetrazol-5-yl	CF_3
99	CF_3	$\text{CH}=\text{NOCH}_3$	CF_3
100	CF_3	$\text{CH}=\text{NOCH}_2\text{CH}=\text{CH}_2$	CF_3
101	CF_3	$\text{CH}=\text{NOCH}_2\text{C}\equiv\text{CH}$	CF_3
102	CF_3	4-(1-methylethoxycarbonyl)phenoxy	CF_3
103	OCF_3	$\text{CH}=\text{NOCH}_2\text{C}\equiv\text{CH}$	OCF_3
104	OCF_3	pyrid-2-yloxy	OCF_3
105	OCF_3	pyrimidin-2-yloxy	OCF_3
106	OCF_3	6-chloro-pyridazin-3yloxy	OCF_3
107	OCF_3	6-methoxy-pyridazin-3yloxy	OCF_3
108	OCF_3	$\text{CH}=\text{NOCH}_3$	OCF_3
109	OCF_3	$\text{CH}=\text{NOCH}_2\text{CH}=\text{CH}_2$	OCF_3
110	OCF_3	4-(1-methylethoxycarbonyl)phenoxy	OCF_3

10

where

m, p, and q are 0; t and u are 1; r is 1, forming an N-oxide; R is phenyl substituted with R^{14} , R^{15} , R^{16} , R^{17} , R^{18} ; X is OH; A is $-\text{CH}_2-$; and R^1 , R^4 , R^5 , R^{15} , R^{18} , R^7 , R^8 , R^{10} , and R^{11} are H:

15



I - C

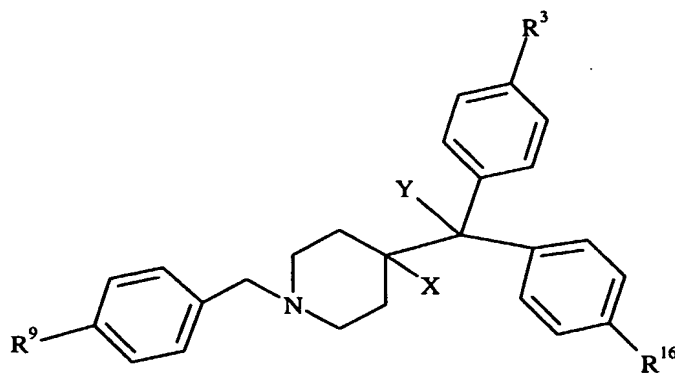
Compd. No.	R ²	R ³	R ⁹	R ¹⁴	R ¹⁶	R ¹⁷	Y
111	H	CF ₃	pyrid-2-yloxy	H	CF ₃	H	H
112	H	OCF ₃	pyrimidin-2-yloxy	H	OCF ₃	H	H
113	H	CF ₃	2-ethyl-2H-tetrazol-5-yl	H	CF ₃	H	H
114	H	OCF ₃	2-ethyl-2H-tetrazol-5-yl	H	OCF ₃	H	H
115	F	Cl	pyrimidin-2-yloxy	H	Cl	F	H
116	H	OCF ₃	pyrimidin-2-yl	H	OCF ₃	H	H
117	H	OCF ₃	pyrid-2-yloxy	H	OCF ₃	H	H
118	H	OCF ₃	CH=NOC ₂ H ₅	H	OCF ₃	H	H
119	H	OCF ₃	NHC(=O)OCH(CH ₃) ₂	H	OCF ₃	H	H
120	H	CF ₃	pyrimidin-2-yloxy	H	CF ₃	H	H
121	H	CF ₃	OC ₃ H ₇	H	CF ₃	H	OH
122	H	OCF ₃	2-methyl-2H-tetrazol-5-yl	H	OCF ₃	H	OH
123	H	OCF ₃	OC ₃ H ₇	H	OCF ₃	H	OH
124	H	CF ₃	CH=NOCH ₃	H	CF ₃	H	H
125	H	CF ₃	CH=NOC ₃ H ₇	H	CF ₃	H	H
126	H	CF ₃	CH=NOCH(CH ₃) ₂	H	CF ₃	H	H
127	H	CF ₃	CH=NOCH ₂ CH=CH ₂	H	CF ₃	H	H
128	H	CF ₃	CH=NOCH ₂ C≡CH	H	CF ₃	H	H
129	H	CF ₃	CH=NOC ₂ H ₅	H	CF ₃	H	H
130	H	OCF ₃	CH=NOCH ₃	H	OCF ₃	H	H
131	H	OCF ₃	CH=NOC ₃ H ₇	H	OCF ₃	H	H
132	H	OCF ₃	CH=NOCH(CH ₃) ₂	H	OCF ₃	H	H
133	H	OCF ₃	CH=NOCH ₂ CH=CH ₂	H	OCF ₃	H	H
134	H	OCF ₃	CH=NOCH ₂ C≡CH	H	OCF ₃	H	H
135	H	OCF ₃	CH=NOC ₂ H ₅	H	OCF ₃	H	H
136	H	CF ₃	CH=NOC ₂ H ₅	H	CF ₃	H	H
137	H	CF ₃	CH=NOC ₂ H ₅	H	CF ₃	H	H
138	H	OCF ₃	phenoxy	H	OCF ₃	H	H

Compd. No.	R ²	R ³	R ⁹	R ¹⁴	R ¹⁶	R ¹⁷	Y
139	H	OCF ₃	4-(1-methylethoxycarbonyl)phenoxy	H	OCF ₃	H	H
140	H	OCF ₃	4-(methoxycarbonyl)phenoxy	H	OCF ₃	H	H
141	H	OCF ₃	6-chloro-pyridazin-3yloxy	H	OCF ₃	H	H
142	H	CF ₃	6-chloro-pyridazin-3yloxy	H	CF ₃	H	H
143	H	OCF ₃	NHCO ₂ CH ₃	H	OCF ₃	H	H
144	H	OCF ₃	NHCO ₂ C ₂ H ₅	H	OCF ₃	H	H
145	H	OCF ₃	NHCO ₂ CH ₂ CH=CH ₂	H	OCF ₃	H	H
146	H	OCF ₃	NHCO ₂ CH ₂ C≡CH	H	OCF ₃	H	H
147	H	CF ₃	NHCO ₂ CH(CH ₃) ₂	H	CF ₃	H	H
148	H	CF ₃	NHCO ₂ CH ₃	H	CF ₃	H	H
149	H	CF ₃	NHCO ₂ CH ₂ CH=CH ₂	H	CF ₃	H	H
150	H	CF ₃	OC(=O)NHCH ₃	H	CF ₃	H	H
151	H	CF ₃	OC(=O)NHC ₂ H ₅	H	CF ₃	H	H
152	H	CF ₃	OC(=O)NHCH(CH ₃) ₂	H	CF ₃	H	H
153	H	CF ₃	OC(=O)NHCH ₂ CH=CH ₂	H	CF ₃	H	H
154	H	OCF ₃	OC(=O)NHCH ₃	H	OCF ₃	H	H
155	H	OCF ₃	OC(=O)NHC ₂ H ₅	H	OCF ₃	H	H
156	H	OCF ₃	OC(=O)NHCH(CH ₃) ₂	H	OCF ₃	H	H
157	H	OCF ₃	OC(=O)NHCH ₂ CH=CH ₂	H	OCF ₃	H	H
158	H	OCF ₃	OC(=O)NHCH ₂ C≡CH	H	OCF ₃	H	H
159	H	CF ₃	OCH ₃	H	CF ₃	H	H
160	H	CF ₃	OC ₂ H ₅	H	CF ₃	H	H
161	H	CF ₃	OC ₃ H ₇	H	CF ₃	H	H
162	H	CF ₃	cyclopentoxo	H	CF ₃	H	H
163	H	OCF ₃	OCH ₃	H	OCF ₃	H	H
164	H	OCF ₃	OC ₂ H ₅	H	OCF ₃	H	H
165	H	OCF ₃	OC ₃ H ₇	H	OCF ₃	H	H
166	H	OCF ₃	OCH(CH ₃) ₂	H	OCF ₃	H	H
167	H	OCF ₃	cyclopentoxo	H	OCF ₃	H	H
168	H	OCHF ₂	pyrid-2-yloxy	H	OCHF ₂	H	H
169	H	OCHF ₂	pyrimidin-2-yloxy	H	OCHF ₂	H	H
170	H	OCHF ₂	CH=NOC ₂ H ₅	H	OCHF ₂	H	H
171	H	OCHF ₂	OC(=O)NHCH(CH ₃) ₂	H	OCHF ₂	H	H
172	H	OCHF ₂	NHCO ₂ CH(CH ₃) ₂	H	OCHF ₂	H	H
173	H	OCHF ₂	OC ₃ H ₇	H	OCHF ₂	H	H
174	H	OCF ₂ CHF ₂	pyrid-2-yloxy	H	OCF ₂ CHF ₂	H	H
175	H	OCF ₂ CHF ₂	pyrimidin-2-yloxy	H	OCF ₂ CHF ₂	H	H
176	H	OCF ₂ CHF ₂	6-chloro-pyridazin-3yloxy	H	OCF ₂ CHF ₂	H	H
177	H	OCF ₂ CHF ₂	OC ₃ H ₇	H	OCF ₂ CHF ₂	H	H
178	H	OCF ₂ CHF ₂	OC(=O)NHCH(CH ₃) ₂	H	OCF ₂ CHF ₂	H	H
179	H	OCF ₂ CHF ₂	NHCO ₂ CH(CH ₃) ₂	H	OCF ₂ CHF ₂	H	H
180	H	OCF ₂ CHF ₂	OCO ₂ CH(CH ₃) ₂	H	OCF ₂ CHF ₂	H	H
181	H	SF ₅	pyrid-2-yloxy	H	SF ₅	H	H

Compd. No.	R ²	R ³	R ⁹	R ¹⁴	R ¹⁶	R ¹⁷	Y
182	H	OCF ₃	pyrid-2-yloxy	Cl	OCF ₃	H	H
183	H	CF ₃	pyrid-2-yloxy	Cl	CF ₃	H	H
184	H	OCF ₃	pyrimidin-2-yloxy	Cl	OCF ₃	H	H
185	H	OCF ₃	CH=NOC ₂ H ₅	Cl	OCF ₃	H	H
186	H	OCF ₃	6-chloro-pyridizin-3yloxy	Cl	OCF ₃	H	H
187	H	OCF ₃	pyrid-2-yloxy	Cl	Cl	H	H
188	H	OCF ₃	pyrid-2-yloxy	F	OCF ₃	H	H
189	H	OCF ₃	pyrimidin-2-yloxy	F	OCF ₃	H	H
190	H	OCF ₃	CH=NOC ₂ H ₅	F	OCF ₃	H	H
191	H	OCF ₃	CO ₂ CH(CH ₃) ₂	H	OCF ₃	H	H

where

- 5 m, p, q and r are 0; t and u are 1; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸, A is -CH₂-; and R¹, R², R⁴, R⁵, R¹⁴, R¹⁵, R¹⁷, R¹⁸, R⁷, R⁸, R¹⁰ and R¹¹ are H:



I - D

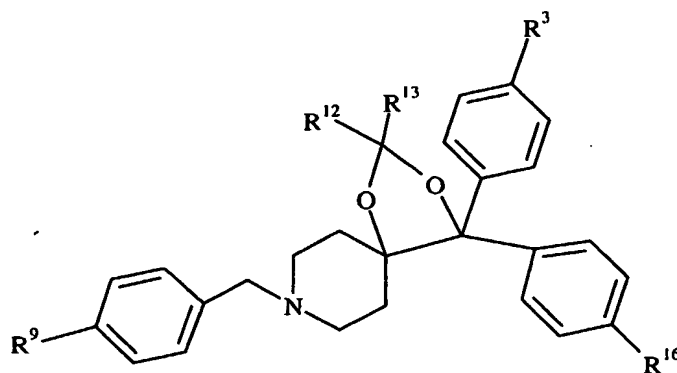
10

Compd. No.	X	Y	R ³	R ⁹	R ¹⁶
192	F	H	OCF ₃	pyrimidin-2-yloxy	OCF ₃
193	F	H	CF ₃	pyrid-2-yloxy	CF ₃
194	F	H	CF ₃	pyrimidin-2-yloxy	CF ₃
195	C(=O)OC ₂ H ₅	Cl	CF ₃	pyrid-2-yloxy	CF ₃
196	C(=O)OC ₂ H ₅	H	CF ₃	pyrid-2-yloxy	CF ₃
197	C(=O)OC ₂ H ₅	F	CF ₃	pyrid-2-yloxy	CF ₃
198	C≡N	Cl	CF ₃	pyrid-2-yloxy	CF ₃
199	OH	CH ₃	CF ₃	pyrid-2-yloxy	CF ₃
200	OH	CH(CH ₃) ₂	CF ₃	pyrid-2-yloxy	CF ₃
201	CH ₂ OH	H	CF ₃	pyrid-2-yloxy	CF ₃
202	OH	CH ₃	CF ₃	pyrimidin-2-yl	CF ₃
203	OH	CH ₃	CF ₃	CH=NOCH ₃	CF ₃
204	OH	CH ₃	OCF ₃	pyrid-2-yl	OCF ₃

Compd. No.	X	Y	R ³	R ⁹	R ¹⁶
205	OH	CH ₃	OCF ₃	pyrimidin-2-yl	OCF ₃
206	OH	CH ₃	OCF ₃	CH=NOCH ₃	OCF ₃
207	C(=O)OCH ₃	CH ₃	OCF ₃	pyrid-2-yl	OCF ₃
208	OSO ₂ CH ₃	CH ₃	OCF ₃	pyrid-2-yl	OCF ₃
209	SH	H	OCF ₃	pyrid-2-yl	OCF ₃
210	C(=O)OH	H	OCF ₃	pyrid-2-yl	OCF ₃

where

m, p, q and r are 0; t and u are 1; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸, R¹, R², R⁴, R⁵, R¹⁴, R¹⁵, R¹⁷, R¹⁸, R⁷, R⁸, R¹⁰ and R¹¹ are H; A is -CH₂-; and X and Y are taken together with -OCR¹²R¹³O-, forming a 1,3-dioxolane ring:

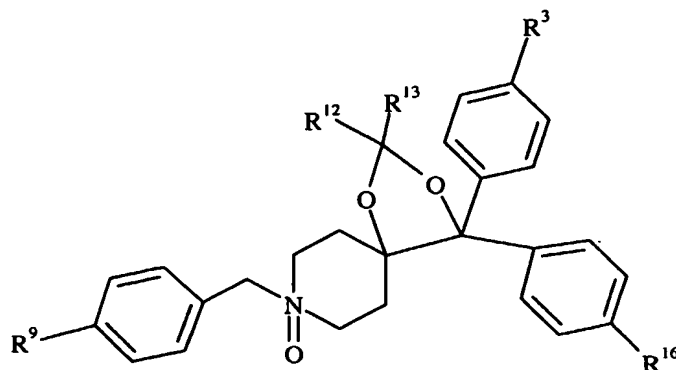


I - E

Compd. No.	R ³	R ⁹	R ¹²	R ¹³	R ¹⁶
211	CF ₃	pyrid-2-yloxy	H	H	CF ₃
212	CF ₃	pyrimidin-2-yloxy	H	H	CF ₃
213	CF ₃	6-chloro-pyridazin-3-yloxy	H	H	CF ₃
214	CF ₃	2-ethyl-2H-tetrazol-5-yl	H	H	CF ₃
215	CF ₃	OC ₃ H ₇	H	H	CF ₃
216	CF ₃	CH=NOCH ₃	H	H	CF ₃
217	CF ₃	CH=NOC ₂ H ₅	H	H	CF ₃
218	CF ₃	pyrid-2-yloxy	CH ₃	CH ₃	CF ₃
219	CF ₃	pyrimidin-2-yloxy	CH ₃	CH ₃	CF ₃
220	CF ₃	6-chloro-pyridazin-3-yloxy	CH ₃	CH ₃	CF ₃
221	CF ₃	2-ethyl-2H-tetrazol-5-yl	CH ₃	CH ₃	CF ₃
222	CF ₃	OC ₃ H ₇	CH ₃	CH ₃	CF ₃
223	CF ₃	CH=NOCH ₃	CH ₃	CH ₃	CF ₃
224	CF ₃	CH=NOC ₂ H ₅	CH ₃	CH ₃	CF ₃
225	CF ₃	OC(=O)NHCH(CH ₃) ₂	CH ₃	CH ₃	CF ₃

where

m, p, and q are 0; t and u are 1; r is 1, forming an N-oxide; R is phenyl substituted with R^{14} , R^{15} , R^{16} , R^{17} , R^{18} ; R^1 , R^2 , R^4 , R^5 , R^{14} , R^{15} , R^{17} , R^{18} , R^7 , R^8 , R^{10} and R^{11} are H; A is $-\text{CH}_2-$; and X and Y are taken together with $-\text{OCR}^{12}\text{R}^{13}\text{O}-$, forming a 1,3-dioxolane ring:



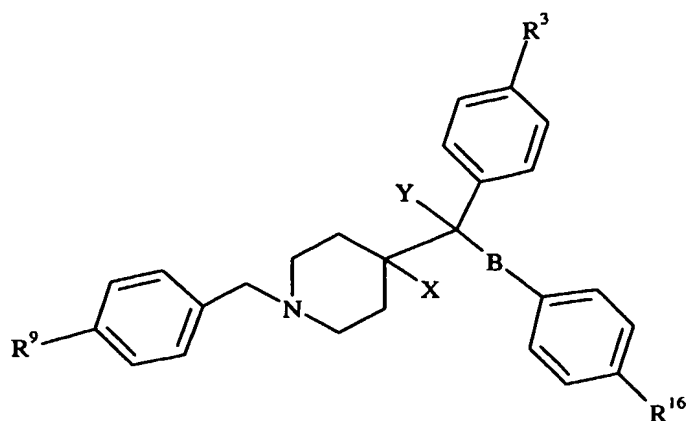
I - F

10

Compd. No.	R^3	R^9	R^{12}	R^{13}	R^{16}
226	CF_3	pyrid-2-yloxy	H	H	CF_3
227	CF_3	pyrimidin-2-yloxy	H	H	CF_3
228	CF_3	6-chloro-pyridizin-3yloxy	H	H	CF_3
229	CF_3	2-ethyl-2H-tetrazol-5-yl	H	H	CF_3
230	CF_3	OC_3H_7	H	H	CF_3
231	CF_3	$\text{CH}=\text{NOCH}_3$	H	H	CF_3
232	CF_3	$\text{CH}=\text{NOC}_2\text{H}_5$	H	H	CF_3
233	CF_3	pyrid-2-yloxy	CH_3	CH_3	CF_3
234	CF_3	pyrimidin-2-yloxy	CH_3	CH_3	CF_3
235	CF_3	6-chloro-pyridizin-3yloxy	CH_3	CH_3	CF_3
236	CF_3	2-ethyl-2H-tetrazol-5-yl	CH_3	CH_3	CF_3
237	CF_3	OC_3H_7	CH_3	CH_3	CF_3
238	CF_3	$\text{CH}=\text{NOCH}_3$	CH_3	CH_3	CF_3
239	CF_3	$\text{CH}=\text{NOC}_2\text{H}_5$	CH_3	CH_3	CF_3
240	CF_3	$\text{OC}(=\text{O})\text{NHCH}(\text{CH}_3)_2$	CH_3	CH_3	CF_3

where

p, q and r are 0; m, t and u are 1; R is phenyl substituted with R^{14} , R^{15} , R^{16} , R^{17} , R^{18} ; A is $-\text{CH}_2-$; and R^1 , R^2 , R^4 , R^5 , R^{14} , R^{15} , R^{17} , R^{18} , R^7 , R^8 , R^{10} and R^{11} are H:



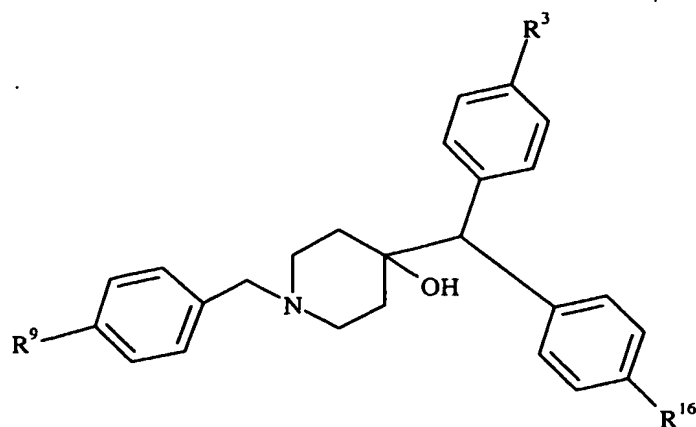
I - G

Compd. No.	R ³	R ⁹	X	Y	B	R ¹⁶
241	CF ₃	pyrid-2-yloxy	OH	H	O	CF ₃
242	CF ₃	pyrid-2-yloxy	OH	CH ₃	O	CF ₃
243	CF ₃	pyrimidin-2-yloxy	OH	H	O	CF ₃
244	CF ₃	CH=NOCH ₃	OH	H	O	CF ₃
245	CF ₃	pyrid-2-yloxy	OH	H	OC(=O)NH	CF ₃
246	OCF ₃	pyrid-2-yloxy	OH	H	O	OCF ₃
247	OCF ₃	pyrid-2-yloxy	OH	CH ₃	O	OCF ₃
248	OCF ₃	pyrid-2-yloxy	F	H	O	OCF ₃
249	OCF ₃	pyrid-2-yloxy	F	CH ₃	O	OCF ₃
250	OCF ₃	pyrid-2-yloxy	F	H	OC(=O)NH	OCF ₃
251	OCF ₃	CH=NOCH ₃	F	H	O	OCF ₃
252	OCF ₃	pyrid-2-yloxy	OH	H	NHSO ₂	OCF ₃
253	OCF ₃	pyrid-2-yloxy	OH	H	OCH ₂	OCF ₃
254	OCF ₃	pyrid-2-yloxy	OH	H	CH ₂ O	OCF ₃

5 where

m, p, q and r are 0; t and u are 1; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸; X is OH; A is -CH₂-; and R¹, R², R⁴, R⁵, R⁷, R⁸, R¹⁰, R¹¹, R¹⁴, R¹⁵, R¹⁷, R¹⁸ and Y are H:

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I - H

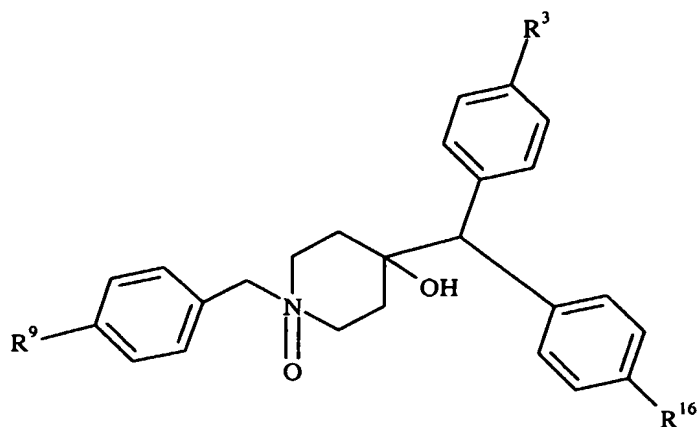
Compd No.	R ³	R ⁹	R ¹⁶
255	OCF ₃	6-methyl-pyridazin-3-yloxy	OCF ₃
256	CHO	pyrid-2-yloxy	OCF ₃
257	C(CH ₃) ₃	pyrid-2-yloxy	OCF ₃
258	OCF ₃	CH=NOCH ₂ CH ₂ OCH ₂ CH ₂ OCH ₃	OCF ₃
259	OCF ₃	CH=NOCH ₂ OCH ₂ CH ₃	OCF ₃
260	OCH ₃	pyrid-2-yloxy	OCF ₃
261	Br	pyrid-2-yloxy	OCF ₃
262	CH ₃	pyrid-2-yloxy	OCF ₃
263	OCF ₃	1,3-dioxolan-2-yl	OCF ₃
264	CF ₃	1,3-dioxolan-2-yl	CF ₃
265	CF ₃	pyrid-2-yloxy	OCF ₃
266	OCF ₃	CH=NOCH ₂ CH ₂ OCH ₃	OCF ₃
267	Cl	pyrid-2-yloxy	Cl
268	F	pyrid-2-yloxy	OCF ₃
269	OCHF ₂	pyrid-2-yloxy	OCF ₃
270	CF ₃	CH=NOH	CF ₃
271	OCF ₃	3-methyl-pyrid-2-yloxy	OCF ₃
272	OCF ₃	5-methyl-pyrid-2-yloxy	OCF ₃
273	OCF ₃	CH(OCH ₂ CH ₃) ₂	OCF ₃
274	OCF ₃	6-methoxy-pyridazin-3-yloxy	OCF ₃
275	OCF ₃	CH=NOCH ₂ C≡N	OCF ₃
276	OCF ₃	6-methyl-pyrid-2-yloxy	OCF ₃
277	OCF ₃	C(Cl)=NOCH ₃	OCF ₃
278	OCF ₃	4-chlorophenylCH(OH)	OCF ₃
279	OCF ₃	OCH ₂ C(=O)OCH ₂ CH ₃	OCF ₃
280	OCF ₃	5,5-dimethyl-1,3-dioxan-2-yl	OCF ₃
281	OCF ₃	3-cyano-pyrid-2-yloxy	OCF ₃
282	OCF ₃	pyrid-2-yl-CH(OH)	OCF ₃
283	OCF ₃	5-cyano-pyrid-2-yloxy	OCF ₃
284	OCF ₃	pyrid-2-yloxy	CH=NOCH ₂ CH=C(CH ₃)CH ₂ - CH ₂ CH=C(CH ₃) ₂
285	OCF ₃	C(=O)NHOCH ₃	OCF ₃
286	OCF ₃	3-chlorophenoxy	OCF ₃
287	OCF ₃	pyrid-2-yloxy	CH(OH)CH ₃
288	H	pyrid-2-yloxy	OCF ₃
289	OCF ₃	4-chlorophenyl-CH(OC(=O)CH ₃)	OCF ₃
290	OCF ₃	4-chlorophenoxy	OCF ₃
291	CF ₃	OCH ₂ C(=O)OCH ₂ CH ₃	CF ₃

Compd No.	R ³	R ⁹	R ¹⁶
292	OCF ₃	1,3-dioxan-2-yl	OCF ₃
293	OCF ₃	OCH ₂ CH=C(Cl) ₂	OCF ₃
294	OCF ₃	3-(dimethoxymethyl)-pyrid-2-yl	OCF ₃
295	1,3-dioxolan-2-yl	pyrid-2-yloxy	OCF ₃
296	OCF ₃	C(C≡N)=NOCH ₃	OCF ₃
297	OCF ₃	2-chlorophenoxy	OCF ₃
298	OCF ₃	OC(=O)NH-cyclopropane	OCF ₃
299	CF ₃	cyclopropylmethoxy	CF ₃
300	OCF ₃	OC(=O)NHC ₃ H ₇	OCF ₃
301	OCF ₃	CH=NOCH ₂ CF ₃	OCF ₃
302	CF ₃	OH	CF ₃
303	CF ₃	OCH(CH ₃) ₂	CF ₃
304	OCF ₃	phenylmethoxy	OCF ₃
305	OCF ₃	CH=NHOCH ₂ CH=C(CH ₃)CH ₂ CH ₂ -CH=C(CH ₃) ₂	OCF ₃
306	OCF ₂ CHFCF ₃	pyrimidin-2-yloxy	OCF ₂ CHFCF ₃
307	CF ₃	6-chloro-pyridazin-3-yloxy	CF ₃
308	OCF ₃	OCH ₂ CH ₂ OCH ₃	OCF ₃
309	OCF ₃	pyrid-2-yl	OCF ₃
310	OCF ₃	NHC(=O)OCH ₂ CH ₂ CH ₃	OCF ₃
311	OCF ₃	OC(=O)NHC(CH ₃) ₃	OCF ₃
312	OCF ₃	CH=NOCH ₂ CH ₂ F	OCF ₃
313	CF ₃	NHC(=O)OCH ₂ CH ₂ CH ₃	CF ₃
314	Cl	pyrid-2-yloxy	OCF ₃
315	OCF ₃	cyclopropylmethoxy	OCF ₃
316	CF ₃	phenylmethoxy	CF ₃
317	OCF ₃	OC(=O)NHCH ₂ C(=O)OC ₂ H ₅	OCF ₃
318	CF ₃	NHC(=O)OCH ₂ CH ₃	CF ₃
319	CF ₃	NHC(=O)OCH ₂ -cyclopropane	CF ₃
320	OCF ₃	OC(=O)NH-cyclopentane	OCF ₃
321	OCF ₃	4-methyl-pyrid-2-yloxy	OCF ₃
322	OCF ₃	3-chloro-pyrid-2-yloxy	OCF ₃
323	OCF ₃	OC(=O)NHCH ₂ (3,4-dichlorophenyl)	OCF ₃
324	OCF ₃	OC(=O)NH(4-chlorophenyl)	OCF ₃
325	Cl	pyrimidin-2-yloxy	OCF ₃
326	OCF ₃	OC(=O)NHC ₄ H ₉	OCF ₃
327	OCF ₃	NHC(=O)OCH ₂ -cyclopropane	OCF ₃
328	Cl	CH=NOC ₂ H ₅	OCF ₃
329	OCF ₃	OC(=O)NH-cyclohexane	OCF ₃
330	CF ₃	C(=O)OCH(CH ₃) ₂	CF ₃
331	CF ₃	OH	CF ₃
332	OCF ₃	NHC(=O)C(CH ₂ CH ₂)C≡N	OCF ₃
333	CF ₃	NHC(=O)C(CH ₂ CH ₂)C≡N	CF ₃

where

- 5 m, p, and q are 0; t and u are 1; r is 1, forming an N-oxide; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸; X is OH; A is -CH₂-; and R¹, R², R⁴, R⁵, R⁷, R⁸, R¹⁰, R¹¹, R¹⁴, R¹⁵, R¹⁷, R¹⁸ and Y are H:

71



I - J

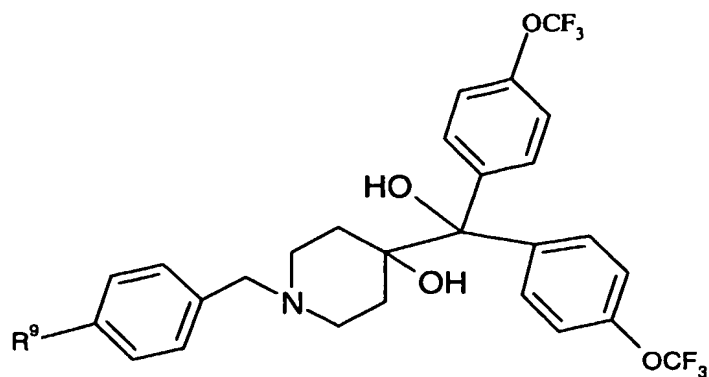
Compd No.	R ³	R ⁹	R ¹⁶
334	CHO	pyrid-2-yloxy	OCF ₃
335	C(CH ₃) ₃	pyrid-2-yloxy	OCF ₃
336	OCF ₃	CH=NOCH ₂ CH ₂ OCH ₂ CH ₂ OCH ₃	OCF ₃
337	OCF ₃	CH=NOCH ₂ OCH ₂ CH ₃	OCF ₃
338	OCH ₃	pyrid-2-yloxy	OCF ₃
339	Br	pyrid-2-yloxy	OCF ₃
340	CH ₃	pyrid-2-yloxy	OCF ₃
341	OCF ₃	1,3-dioxolan-2-yl	OCF ₃
342	CF ₃	1,3-dioxolan-2-yl	CF ₃
343	CF ₃	pyrid-2-yloxy	OCF ₃
344	OCF ₃	CH=NOCH ₂ CH ₂ OCH ₃	OCF ₃
345	Cl	pyrid-2-yloxy	Cl
346	F	pyrid-2-yloxy	OCF ₃
347	OCHF ₂	pyrid-2-yloxy	OCF ₃
348	CF ₃	CH=NOH	CF ₃
349	OCF ₃	3-methyl-pyrid-2-yloxy	OCF ₃
350	OCF ₃	5-methyl-pyrid-2-yloxy	OCF ₃
351	OCF ₃	CH(OCH ₂ CH ₃) ₂	OCF ₃
352	OCF ₃	6-methoxy-pyridazin-3-yloxy	OCF ₃
353	OCF ₃	CH=NOCH ₂ C≡N	OCF ₃
354	OCF ₃	6-methyl-pyrid-2-yloxy	OCF ₃
355	OCF ₃	C(Cl)=NOCH ₃	OCF ₃
356	OCF ₃	4-chlorophenylCH(OH)	OCF ₃
357	OCF ₃	OCH ₂ C(=O)OCH ₂ CH ₃	OCF ₃
358	OCF ₃	5,5-dimethyl-1,3-dioxan-2-yl	OCF ₃
359	OCF ₃	3-cyano-pyrid-2-yloxy	OCF ₃
360	OCF ₃	pyrid-2-yl-CH(OH)	OCF ₃
361	OCF ₃	5-cyano-pyrid-2-yloxy	OCF ₃
362	OCF ₃	pyrid-2-yloxy	CH=NOCH ₂ CH=C(CH ₃)CH ₂ - CH ₂ CH=C(CH ₃) ₂
363	OCF ₃	C(=O)NHOCH ₃	OCF ₃
364	OCF ₃	3-chlorophenoxy	OCF ₃
365	OCF ₃	pyrid-2-yloxy	CH(OH)CH ₃
366	H	pyrid-2-yloxy	OCF ₃
367	OCF ₃	4-chlorophenyl-CH(OC(=O)CH ₃)	OCF ₃
368	OCF ₃	4-chlorophenoxy	OCF ₃
369	CF ₃	OCH ₂ C(=O)OCH ₂ CH ₃	CF ₃
370	OCF ₃	1,3-dioxan-2-yl	OCF ₃

Compd No.	R ³	R ⁹	R ¹⁶
371	OCF ₃	OCH ₂ CH=C(Cl) ₂	OCF ₃
372	OCF ₃	3-(dimethoxymethyl)-pyrid-2-yl	OCF ₃
373	1,3-dioxolan-2-yl	pyrid-2-yloxy	OCF ₃
374	OCF ₃	C(C≡N)=NOCH ₃	OCF ₃
375	OCF ₃	2-chlorophenoxy	OCF ₃
376	OCF ₃	OC(=O)NH-cyclopropane	OCF ₃
377	OCF ₃	OC(=O)NHC ₃ H ₇	OCF ₃
378	OCF ₃	CH=NOCH ₂ CF ₃	OCF ₃
379	CF ₃	OH	CF ₃
380	CF ₃	OCH(CH ₃) ₂	CF ₃
381	OCF ₃	phenylmethoxy	OCF ₃
382	OCF ₃	CH=NHOCH ₂ CH=C(CH ₃)CH ₂ CH ₂ -CH=C(CH ₃) ₂	OCF ₃
383	CF ₃	cyclopropylmethoxy	CF ₃
384	CF ₃	6-chloro-pyridazin-3-yloxy	CF ₃
385	OCF ₃	OCH ₂ CH ₂ OCH ₃	OCF ₃
386	OCF ₃	pyrid-2-yl	OCF ₃
387	OCF ₃	NHC(=O)OCH ₂ CH ₂ CH ₃	OCF ₃
388	OCF ₃	OC(=O)NHC(CH ₃) ₃	OCF ₃
389	OCF ₃	CH=NOCH ₂ CH ₂ F	OCF ₃
390	CF ₃	NHC(=O)OCH ₂ CH ₂ CH ₃	CF ₃
391	Cl	pyrid-2-yloxy	OCF ₃
392	OCF ₃	cyclopropylmethoxy	OCF ₃
393	CF ₃	phenylmethoxy	CF ₃
394	OCF ₃	OC(=O)NHCH ₂ C(=O)OC ₂ H ₅	OCF ₃
395	CF ₃	NHC(=O)OCH ₂ CH ₃	CF ₃
396	CF ₃	NHC(=O)OCH ₂ -cyclopropane	CF ₃
397	OCF ₃	OC(=O)NH-cyclopentane	OCF ₃
398	OCF ₃	4-methyl-pyrid-2-yloxy	OCF ₃
399	OCF ₃	3-chloro-pyrid-2-yloxy	OCF ₃
400	OCF ₃	OC(=O)NHCH ₂ (3,4-dichlorophenyl)	OCF ₃
401	OCF ₃	OC(=O)NH(4-chlorophenyl)	OCF ₃
402	Cl	pyrimidin-2-yloxy	OCF ₃
403	OCF ₃	OC(=O)NHC ₄ H ₉	OCF ₃
404	OCF ₃	NHC(=O)OCH ₂ -cyclopropane	OCF ₃
405	Cl	CH=NOC ₂ H ₅	OCF ₃
406	OCF ₃	OC(=O)NH-cyclohexane	OCF ₃
407	CF ₃	C(=O)OCH(CH ₃) ₂	CF ₃
408	CF ₃	OH	CF ₃
409	OCF ₂ CHF _{CF₃}	pyrimidin-2-yloxy	OCF ₂ CHF _{CF₃}
410	OCF ₃	CH=NOCH ₂ C(=O)NH ₂	OCF ₃
411	OCF ₃	3-chloro-pyrid-2-yl	OCF ₃
412	OCF ₃	6-chloro-pyridazin-3-yloxy	OCF ₃
413	OCF ₃	6-methyl-pyridazin-3-yloxy	OCF ₃
414	OCF ₃	NHC(=O)C(CH ₂ CH ₂)C≡N	OCF ₃
415	CF ₃	NHC(=O)C(CH ₂ CH ₂)C≡N	CF ₃

where

m, p, q and r are 0; t and u are 1; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸, X and Y are OH; R³ and R¹⁶ are OCF₃; A is -CH₂-; and R¹, R², R⁴, R⁵, R¹⁴, R¹⁵, R¹⁷, R¹⁸, R⁷, R⁸, R¹⁰, and R¹¹ are H:

73



I - K

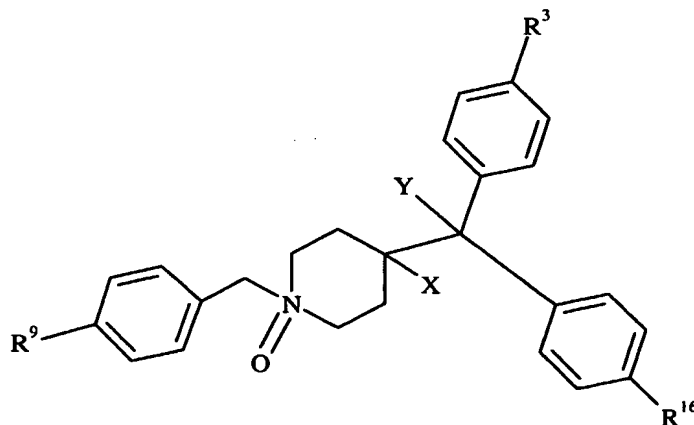
Compd. No.	R ⁹
416	2-ethyl-2H-tetrazol-5-yl
417	CH=NOCH ₂ CH ₃
418	CHO
419	NHC(=O)OCH(CH ₃) ₂
420	NHC(=O)C(CH ₂ CH ₂)C≡N

5

where

m, p, and q are 0; t and u are 1; r is 1, forming an N-oxide; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸; A is -CH₂-; and R¹, R², R⁴, R⁵, R¹⁴, R¹⁵, R¹⁷, R¹⁸, R⁷, R⁸, R¹⁰, and R¹¹ are H:

10



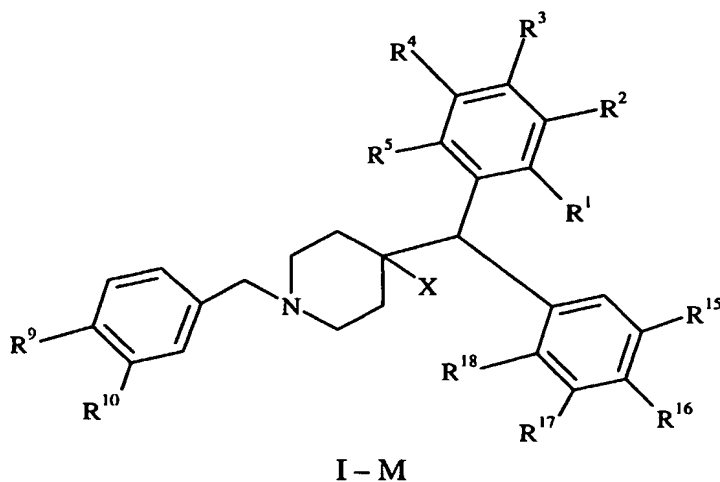
I - L

Compd. No.	R ³	R ⁹	R ¹⁶	X	Y
421	OCF ₃	pyrid-2-yloxy	OCF ₃	OH	OH
422	OCF ₃	pyrimidin-2-yloxy	OCF ₃	OH	OH
423	OCF ₃	NHC(=O)OCH(CH ₃) ₂	OCF ₃	OH	OH
424	OCF ₃	CH=NOCH ₂ CH ₃	OCF ₃	OH	OH

Compd. No.	R ³	R ⁹	R ¹⁶	X	Y
425	OCF ₃	6-chloro-pyridazin-3-yloxy	OCF ₃	OH	OH
426	OCF ₃	2-ethyl-2H-tetrazol-5-yl	OCF ₃	OH	OH
428	OCF ₃	NHC(=O)C(CH ₂ CH ₂)C≡N	OCF ₃	OH	OH
429	CF ₃	NHC(=O)C(CH ₂ CH ₂)C≡N	CF ₃	OH	OH
430	OCF ₃	pyrid-2-yloxy	OCF ₃	OH	F
431	OCF ₃	pyrid-2-yloxy	OCF ₃	X and Y taken together with bridging group -O-	
432	OCF ₃	pyrimidin-2-yloxy	OCF ₃	X and Y taken together with bridging group -O-	

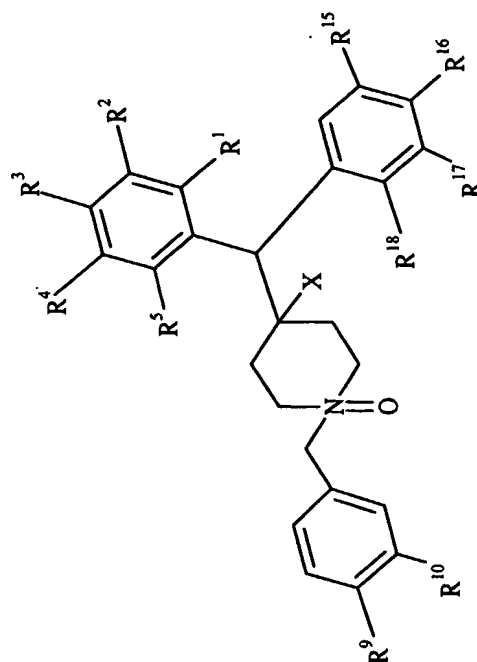
where

- 5 m, p, q and r are 0; t and u are 1; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸; A is -CH₂-; and R⁷, R⁸, R¹¹, R¹⁴ and Y are H:



Compd No.	R ¹	R ²	R ³	R ⁴	R ⁵	R ⁹	R ¹⁰	R ¹⁵	R ¹⁶	R ¹⁷	R ¹⁸	X
433	Cl	H	Cl	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
434	F	H	H	H	H	pyrimidin-2-yloxy	H	H	H	H	F	OH
435	H	CH ₃	Cl	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
436	CH ₃	H	Cl	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
437	Cl	H	H	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
438	Cl	Cl	H	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
439	H	Cl	H	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
440	H	H	CH=NO- C ₂ H ₅	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
441	H	Cl	Cl	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
442	H	Cl	H	Cl	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
443	Cl	H	H	Cl	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
444	Cl	H	Cl	H	H	CH=NOC ₂ H ₅	H	H	OCF ₃	H	H	OH
445	F	H	Cl	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
446	H	H	R ³ and R ⁴ taken together with bridging group - OC(F ₂)O-	H	H	pyrid-2-yloxy	H	H	R ¹⁶ and R ¹⁷ taken together with bridging group -OC(F ₂)O-	H	H	OH
447	H	H	OCF ₃	H	H	pyrid-2-yloxy	OCH ₃	H	OCF ₃	H	H	OH
448	H	H	CF ₃	H	H	pyrimidin-2-yloxy	H	H	CF ₃	H	H	CH ₂ OH
449	H	H	CF ₃	H	H	OCH ₂ -cyclopropane	H	H	CF ₃	H	H	CH ₂ N=N
450	H	H	CF ₃	H	H	OCH ₂ -cyclopropane	H	H	CF ₃	H	H	CH ₂ NH ₂
451	H	H	CF ₃	H	H	pyrimidin-2-yloxy	H	H	CF ₃	H	H	CH ₂ NH-
452	H	H	CF ₃	H	H	pyrimidin-2-yloxy	H	H	CF ₃	H	H	C(=O)-CH ₃ CH ₂ O-C(=O)- CH ₃

m, p, and q are 0; t and u are 1; r is 1, forming an N-oxide; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸; A is -CH₂-; and R⁷, R⁸, R¹¹, R¹⁴ and Y are H:



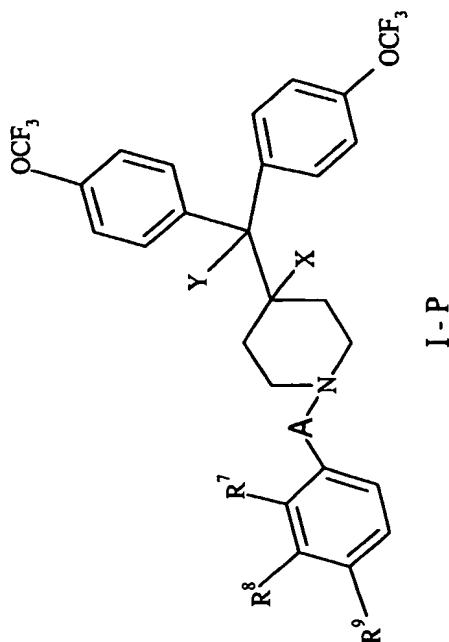
I - N

Compd No.	R ¹	R ²	R ³	R ⁴	R ⁵	R ⁹	R ¹⁰	R ¹⁵	R ¹⁶	R ¹⁷	R ¹⁸	X
453	Cl	H	Cl	H	H	CH=NOC ₂ H ₅	H	H	OCF ₃	H	H	OH
454	Cl	H	Cl	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
455	H	Cl	Cl	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
456	Cl	Cl	H	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
457	H	Cl	H	Cl	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
458	H	Cl	H	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
459	Cl	H	H	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
460	Cl	H	H	Cl	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
461	CH ₃	H	Cl	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
462	H	CH ₃	Cl	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH

Compd No.	R ¹	R ²	R ³	R ⁴	R ⁵	R ⁹	R ¹⁰	R ¹⁵	R ¹⁶	R ¹⁷	R ¹⁸	X
463	F	H	Cl	H	H	pyrid-2-yloxy	H	H	OCF ₃	H	H	OH
464	H	R ² and R ³ taken together with bridging group -OC(F ₂)O-		H	H	pyrid-2-yloxy	H	R ¹⁵ and R ¹⁶ taken together with bridging group -OC(F ₂)O-		H	H	OH
465	H	H	OCF ₃	H	H	pyrid-2-yloxy	OCH ₃	H	OCF ₃	H	H	OH
466	H	H	CF ₃	H	H	pyrimidin-2-yloxy	H	H	CF ₃	H	H	CH ₂ NH- C(=O)-CH ₃
467	H	H	CF ₃	H	H	OCH ₂ -cyclopropane	H	H	CF ₃	H	H	CH ₂ N=N- CH ₂ O-C(=O)- CH ₃
468	H	H	CF ₃	H	H	pyrimidin-2-yloxy	H	H	CF ₃	H	H	

where

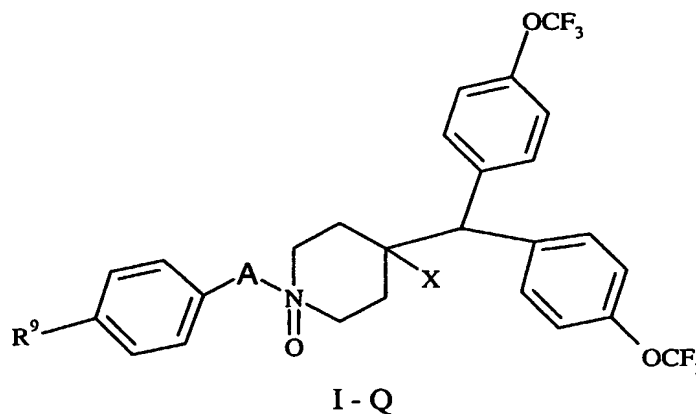
m, p, q and r are 0; t and u are 1; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸, R³ and R¹⁶ are OCF₃; and R¹, R², R⁴, R⁵, R¹⁴, R¹⁵, R¹⁷, R¹⁸, R¹⁰ and R¹¹ are H:



Compd. No.	X	Y	A	R ⁷	R ⁸	R ⁹
470	OH	H	OCH ₂ CH(OH)CH ₂	H	H	Cl
471	OH	H	OCH ₂ CH(OH)CH ₂	H	Cl	H
472	OH	H	OCH ₂ CH(OH)CH ₂	Cl	H	H
473	OH	H	OCH ₂ CH ₂ CH ₂	H	H	Cl
474	OH	H	CH ₂ CH ₂	H	H	Cl
475	OH	H	OCH ₂ CH ₂	H	H	Cl
476	OH	H	OCH ₂ CH ₂ CH ₂ CH ₂	H	H	Cl
477	OH	H	OCH ₂ CH ₂	H	H	Br

where

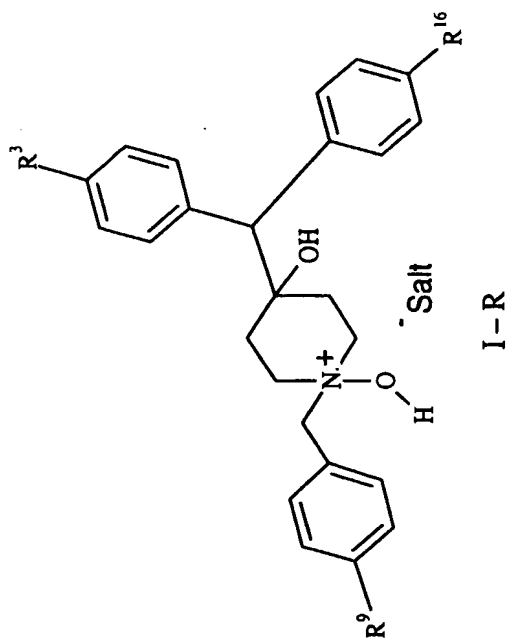
5 m, p and q are 0; t and u are 1; r is 1, forming an N-oxide; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸, R³ and R¹⁶ are OCF₃; and R¹, R², R⁴, R⁵, R⁷, R⁸, R¹⁰, R¹¹, R¹⁴, R¹⁵, R¹⁷, R¹⁸ and Y are H:



Compd. No.	X	A	R ⁹
478	OH	OCH ₂ CH ₂	Cl
479	OH	CH ₂ CH ₂	Cl
480	OH	OCH ₂ CH ₂ CH ₂ CH ₂	Cl
481	OH	OCH ₂ CH ₂ CH ₂	Cl
482	OH	OCH ₂ CH ₂	Br

where.

15 m, p, and q are 0; r, t and u are 1; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸; X is OH; A is -CH₂-; and R¹, R², R⁴, R⁵, R⁷, R⁸, R¹⁰, R¹¹, R¹⁴, R¹⁵, R¹⁷, R¹⁸ and Y are H:



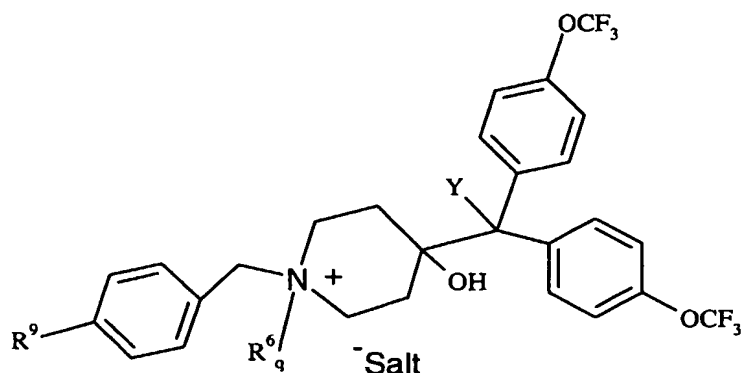
Compd No.	R ³	R ⁹	R ¹⁶	Salt
483	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	3-hydroxypropanesulfonic acid
484	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	2-ethoxypropanoic acid
485	OCF ₃	6-chloro-pyridazin-3-ylloxy	OCF ₃	2-ethylhexanoic acid
486	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	2-ethylhexanoic acid
487	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	1,1,2,2,3,3,4,4,4-nonafluorobutanesulfonic acid
488	CF ₃	6-chloro-pyridazin-3-ylloxy	CF ₃	2-hydroxypropane-1,2,3-tricarboxylic acid
489	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	2-hydroxypropane-1,2,3-tricarboxylic acid
490	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	((4R)-7,7-dimethyl-2-oxobicyclo[2.2.1]heptyl)methanesulfonic acid
491	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	ethanesulfonic acid
492	OCF ₃	6-chloro-pyridazin-3-ylloxy	OCF ₃	2-hydroxypropane-1,2,3-tricarboxylic acid
493	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	cyclohexanecarboxylic acid
494	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	2-hydroxypropane-1,2,3-tricarboxylic acid
495	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	1,1,2,2,3,3,4,4,5,5,6,6,7,7,8,8-heptafluorooctanesulfonic acid

Compd No.	R ³	R ⁹	R ¹⁶	Salt
496	OCF ₃	pyrid-2-yloxy	OCF ₃	1,1,2,2,3,3,4,4,5,5,6,6,7,7,8,8-heptafluorooctanesulfonic acid
497	OCF ₃	pyrid-2-yloxy	OCF ₃	Cl
498	OCF ₃	pyrid-2-yloxy	OCF ₃	((4S)-7,7-dimethyl-2-oxobicyclo[2.2.1]heptyl)methanesulfonic acid
499	OCF ₃	pyrid-2-yloxy	OCF ₃	2-ethylhexanoic acid
500	OCF ₃	pyrid-2-yloxy	OCF ₃	3-hydroxypropanesulfonic acid
501	OCF ₃	CH(Cl)=NOCH ₃	OCF ₃	2-hydroxypropane-1,2,3-tricarboxylic acid
502	OCF ₃	pyrid-2-yloxy	OCF ₃	cyclohexanecarboxylic acid
503	OCF ₃	pyrid-2-yloxy	OCF ₃	2-ethoxypropanoic acid
504	OCF ₃	pyrid-2-yloxy	OCF ₃	2-hydroxypropane-1,2,3-tricarboxylic acid
505	OCF ₃	pyrid-2-yloxy	OCF ₃	((4R)-7,7-dimethyl-2-oxobicyclo[2.2.1]heptyl)methanesulfonic acid
506	OCF ₃	pyrid-2-yloxy	OCF ₃	ethanesulfonic acid
507	OCF ₃	CH=NOC ₂ H ₅	OCF ₃	((4R)-7,7-dimethyl-2-oxobicyclo[2.2.1]heptyl)methanesulfonic acid
508	OCF ₃	pyrid-2-yloxy	OCF ₃	1,1,2,2,3,3,4,4,4-nonafluorobutanesulfonic acid
509	OCF ₃	pyrid-2-yloxy	OCF ₃	3,7-dimethyloct-6-enoic acid
510	OCF ₃	pyrid-2-yloxy	OCF ₃	2-hydroxyacetic acid
511	OCF ₃	pyrid-2-yloxy	OCF ₃	adamantanecarboxylic acid
512	OCF ₃	pyrid-2-yloxy	OCF ₃	pentanedioic acid
513	OCF ₃	pyrid-2-yloxy	OCF ₃	(9E)octadec-9-enoic acid
514	OCF ₃	pyrid-2-yloxy	OCF ₃	heptanedioic acid
515	OCF ₃	pyrid-2-yloxy	OCF ₃	4-dodecylbenzenesulfonic acid
516	OCF ₃	pyrid-2-yloxy	OCF ₃	hexanedioic acid
517	OCF ₃	pyrid-2-yloxy	OCF ₃	octanoic acid
518	CF ₃	6-chloro-pyridazin-3-yloxy	CF ₃	ethanesulfonic acid
519	CF ₃	6-chloro-pyridazin-3-yloxy	CF ₃	3-((1Z)-2-chloro-3,3,3-trifluoroprop-1-enyl)-2,2-dimethylcyclopropanecarboxylic acid
520	CF ₃	6-chloro-pyridazin-3-yloxy	CF ₃	1,1,2,2,3,3,4,4,5,5,6,6,7,7,8,8,9,9,9-nonadecafluorononanesulfonic acid

where

m, p and r are 0; t and u are 1; R is phenyl substituted with R^{14} , R^{15} , R^{16} , R^{17} , R^{18} ; X is OH; A is $-\text{CH}_2-$; R^3 and R^{16} are OCF_3 ; and R^1 , R^2 , R^4 , R^5 , R^7 , R^8 , R^{10} , R^{11} , R^{14} ,

5 R^{15} , R^{17} and R^{18} are H:



I - S

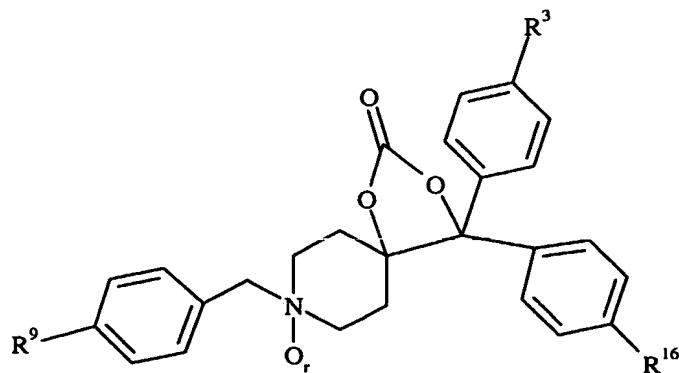
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Compd No.	q	Y	R^6	R^9	Salt
521	1	OH	(methylethoxy)-carboxamide-phenyl-4-meth-yl	$\text{NHC}(=\text{O})\text{OCH}(\text{CH}_3)_2$	Cl^-
522	0	H	-	pyrid-2-yloxy	$\text{H}^+ \cdot \text{Cl}^-$

15 where

m, p and q are 0; t and u are 1; R is phenyl substituted with R^{14} , R^{15} , R^{16} , R^{17} , R^{18} ; R^1 , R^2 , R^4 , R^5 , R^{14} , R^{15} , R^{17} , R^{18} , R^7 , R^8 , R^{10} and R^{11} are H; A is $-\text{CH}_2-$; and X and Y are taken together with $-\text{OC}(=\text{O})\text{O}-$, forming a 1,3-dioxol-2-one ring:

20



I - T

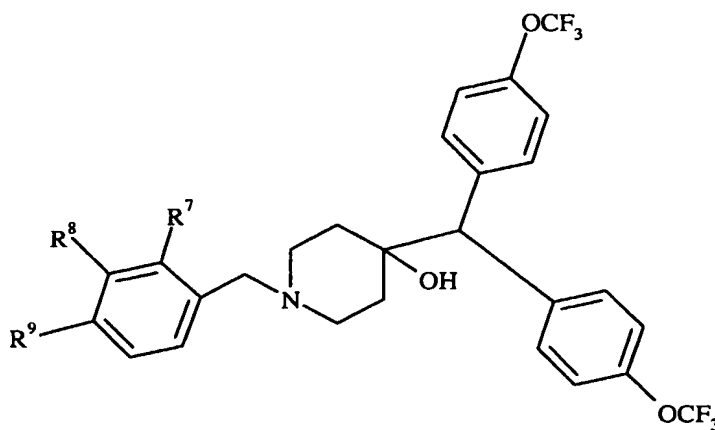
note if $r=1$, the n-oxide is formed:

Compd. No.	r	R ³	R ⁹	R ¹⁶
523	0	CF ₃	pyrid-2-yloxy	CF ₃
524	0	CF ₃	pyrimidin-2-yloxy	CF ₃
525	0	CF ₃	6-chloro-pyridazin-3yloxy	CF ₃
526	0	CF ₃	2-ethyl-2H-tetrazol-5-yl	CF ₃
527	0	CF ₃	OC ₃ H ₇	CF ₃
528	0	CF ₃	CH=NOCH ₃	CF ₃
529	0	CF ₃	CH=NOC ₂ H ₅	CF ₃
530	0	CF ₃	pyrid-2-yloxy	CF ₃
531	0	CF ₃	pyrimidin-2-yloxy	CF ₃
532	0	CF ₃	6-chloro-pyridazin-3yloxy	CF ₃
533	0	CF ₃	2-ethyl-2H-tetrazol-5-yl	CF ₃
534	0	CF ₃	OC ₃ H ₇	CF ₃
535	0	CF ₃	CH=NOCH ₃	CF ₃
536	0	CF ₃	CH=NOC ₂ H ₅	CF ₃
537	0	CF ₃	OC(=O)NHCH(CH ₃) ₂	CF ₃
538	1	CF ₃	pyrid-2-yloxy	CF ₃
539	1	CF ₃	pyrimidin-2-yloxy	CF ₃
540	1	CF ₃	6-chloro-pyridazin-3yloxy	CF ₃
541	1	CF ₃	2-ethyl-2H-tetrazol-5-yl	CF ₃
542	1	CF ₃	OC ₃ H ₇	CF ₃
543	1	CF ₃	CH=NOCH ₃	CF ₃
544	1	CF ₃	CH=NOC ₂ H ₅	CF ₃
545	1	CF ₃	pyrid-2-yloxy	CF ₃
546	1	CF ₃	pyrimidin-2-yloxy	CF ₃
547	1	CF ₃	6-chloro-pyridazin-3yloxy	CF ₃
548	1	CF ₃	2-ethyl-2H-tetrazol-5-yl	CF ₃
549	1	CF ₃	OC ₃ H ₇	CF ₃
550	1	CF ₃	CH=NOCH ₃	CF ₃
551	1	CF ₃	CH=NOC ₂ H ₅	CF ₃
552	1	CF ₃	OC(=O)NHCH(CH ₃) ₂	CF ₃
553	0	OCF ₃	pyrid-2-yloxy	OCF ₃
554	0	OCF ₃	pyrimidin-2-yloxy	OCF ₃
555	0	OCF ₃	6-chloro-pyridazin-3yloxy	OCF ₃
556	0	OCF ₃	2-ethyl-2H-tetrazol-5-yl	OCF ₃
557	0	OCF ₃	OC ₃ H ₇	OCF ₃
558	0	OCF ₃	CH=NOCH ₃	OCF ₃
559	0	OCF ₃	CH=NOC ₂ H ₅	OCF ₃
560	0	OCF ₃	pyrid-2-yloxy	OCF ₃
561	0	OCF ₃	pyrimidin-2-yloxy	OCF ₃
562	0	OCF ₃	6-chloro-pyridazin-3yloxy	OCF ₃

Compd. No.	r	R ³	R ⁹	R ¹⁶
563	0	OCF ₃	2-ethyl-2H-tetrazol-5-yl	OCF ₃
564	0	OCF ₃	OC ₃ H ₇	OCF ₃
565	0	OCF ₃	CH=NOCH ₃	OCF ₃
566	0	OCF ₃	CH=NOC ₂ H ₅	OCF ₃
567	0	OCF ₃	OC(=O)NHCH(CH ₃) ₂	OCF ₃
568	1	OCF ₃	pyrid-2-yloxy	OCF ₃
569	1	OCF ₃	pyrimidin-2-yloxy	OCF ₃
570	1	OCF ₃	6-chloro-pyridazin-3yloxy	OCF ₃
571	1	OCF ₃	2-ethyl-2H-tetrazol-5-yl	OCF ₃
572	1	OCF ₃	OC ₃ H ₇	OCF ₃
573	1	OCF ₃	CH=NOCH ₃	OCF ₃
574	1	OCF ₃	CH=NOC ₂ H ₅	OCF ₃
575	1	OCF ₃	pyrid-2-yloxy	OCF ₃
576	1	OCF ₃	pyrimidin-2-yloxy	OCF ₃
577	1	OCF ₃	6-chloro-pyridazin-3yloxy	OCF ₃
578	1	OCF ₃	2-ethyl-2H-tetrazol-5-yl	OCF ₃
579	1	OCF ₃	OC ₃ H ₇	OCF ₃
580	1	OCF ₃	CH=NOCH ₃	OCF ₃
581	1	OCF ₃	CH=NOC ₂ H ₅	OCF ₃
582	1	OCF ₃	OC(=O)NHCH(CH ₃) ₂	OCF ₃

where

m, p, q and r are 0; t and u are 1; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸.
 X is OH; A is -CH₂-; R³ and R¹⁶ are OCF₃; and R¹, R², R⁴, R⁵, R¹⁰, R¹¹, R¹⁴, R¹⁵, R¹⁷,
 5 R¹⁸ and Y are H:



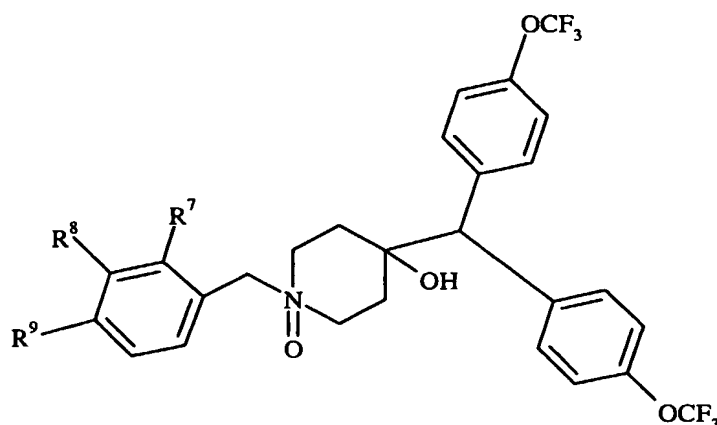
I - U

Compd No.	R ⁷	R ⁸	R ⁹
583	H	H	OCH ₂ CH ₂ OCH ₂ CH ₂ OCH ₃
584	H	H	OCH ₂ CH ₂ CH ₂ OCH ₃
585	H	H	6-methoxy-pyrid-2-yloxy

Compd No.	R ⁷	R ⁸	R ⁹
586	H	methoxy	OCH ₂ CH ₂ OCH ₃
587	H	Cl	OCH ₂ CH ₂ OCH ₃
588	Cl	H	OCH ₂ CH ₂ OCH ₃

where

5 m, p, and q are 0; t and u are 1; r is 1, forming an N-oxide; R is phenyl substituted with R¹⁴, R¹⁵, R¹⁶, R¹⁷, R¹⁸; X is OH; A is -CH₂-; R³ and R¹⁶ are OCF₃; and R¹, R², R⁴, R⁵, R¹⁰, R¹¹, R¹⁴, R¹⁵, R¹⁷, R¹⁸ and Y are H:



I - V

Compd No.	R ⁷	R ⁸	R ⁹
589	H	H	OCH ₂ CH ₂ OCH ₂ CH ₂ OCH ₃
590	H	H	OCH ₂ CH ₂ CH ₂ OCH ₃
591	H	H	6-methoxy-pyrid-2-yloxy
592	H	methoxy	OCH ₂ CH ₂ OCH ₃
593	H	Cl	OCH ₂ CH ₂ OCH ₃
594	Cl	H	OCH ₂ CH ₂ OCH ₃

The following table sets forth physical characterizing data for compounds of formula I:

Table 2 - Physical Characteristics

Cmpd No.	Emperical Formula	Physical State / Melting Point (°C)
1	C ₂₉ H ₂₇ Cl ₂ N ₃ O ₂	Solid Foam
2	C ₃₁ H ₂₇ F ₆ N ₃ O ₃	Solid
3	C ₃₀ H ₂₆ Cl ₂ F ₃ N ₃ O ₂	--
4	C ₂₉ H ₂₅ Cl ₂ F ₂ N ₃ O ₂	Solid Foam
5	C ₃₂ H ₃₂ F ₃ N ₃ O ₄	Solid
6	C ₂₉ H ₂₅ C ₁₄ N ₃ O ₂	Solid Foam
7	C ₃₀ H ₂₇ ClF ₃ N ₃ O ₂	Solid
8	C ₂₉ H ₂₇ F ₂ N ₃ O ₂	--
9	C ₃₀ H ₂₆ Cl ₂ F ₃ N ₃ O ₂	--

Cmpd No.	Emperical Formula	Physical State / Melting Point (°C)
10	$C_{29}H_{25}F_4N_3O_2$	--
11	$C_{30}H_{26}F_5N_3O_2$	--
12	$C_{30}H_{27}F_4N_3O_2$	Solid
13	$C_{30}H_{26}Cl_2F_3N_3O_2$	Solid
14	$C_{30}H_{27}ClF_3N_3O_2$	Solid
15	$C_{30}H_{26}Cl_2F_3N_3O_2$	--
16	$C_{31}H_{27}F_6N_3O_2$	--
17	$C_{30}H_{26}F_5N_3O_2$	Oil
18	$C_{30}H_{27}ClF_3N_3O_2$	Solid
19	$C_{31}H_{27}F_6N_3O_4$	Solid, 68-79
20	$C_{31}H_{27}F_6N_3O_3$	Solid, 59-65
21	$C_{32}H_{28}F_6N_2O_4$	--
22	$C_{32}H_{28}F_6N_2O_2$	Solid
23	$C_{28}H_{29}Cl_2N_5O$	--
24	$C_{28}H_{29}Cl_2N_5O$	Solid
25	$C_{30}H_{29}F_6N_5O_3$	Solid / Solid Foam
26	$C_{28}H_{27}Cl_4N_5O$	Solid Foam
27	$C_{30}H_{29}F_6N_5O$	Solid Foam
28	$C_{30}H_{30}F_6N_2O_4$	Liquid
29	$C_{31}H_{32}F_6N_2O_5$	Solid, 82-85
30	$C_{33}H_{27}F_9N_2O_4$	--
91	$C_{30}H_{31}F_6NO_5$	Solid, 52-57
92	$C_{30}H_{31}F_6NO_3$	Solid, 65-70
93	$C_{29}H_{27}F_6N_5O_4$	Solid
111	$C_{32}H_{28}F_6N_2O_3$	Solid
112	$C_{31}H_{27}F_6N_3O_5$	Solid, 198-202
113	$C_{30}H_{29}F_6N_5O_2$	--
114	$C_{30}H_{29}F_6N_5O_4$	Solid
115	$C_{29}H_{25}Cl_2F_2N_3O_3$	Solid Foam
116	$C_{31}H_{27}F_6N_3O_4$	Solid, 213-215
117	$C_{32}H_{28}F_6N_2O_5$	Solid
118	$C_{30}H_{30}F_6N_2O_5$	Solid, 82-184
119	$C_{31}H_{32}F_6N_2O_6$	Solid, 149-162
120	$C_{31}H_{27}F_6N_3O_3$	Solid
121	$C_{30}H_{31}F_6NO_4$	Solid, 151-154
122	$C_{29}H_{27}F_6N_5O_5$	Solid, 208-211
123	$C_{30}H_{31}F_6NO_6$	Solid, 219-221
192	$C_{31}H_{26}F_7N_3O_3$	Solid
193	$C_{32}H_{27}F_7N_2O$	Solid
194	$C_{31}H_{26}F_7N_3O$	Solid
195	$C_{35}H_{31}ClF_6N_2O_3$	Solid
255	$C_{32}H_{29}F_6N_3O_4$	Solid
256	$C_{32}H_{29}F_3N_2O_4$	Solid Foam, 54-57
257	$C_{35}H_{37}F_3N_2O_3$	Solid Foam, 71-75
258	$C_{33}H_{36}F_6N_2O_6$	Liquid
259	$C_{31}H_{32}F_6N_2O_5$	Liquid
260	$C_{32}H_{31}F_3N_2O_4$	Solid Foam, 54-56
261	$C_{31}H_{28}BrF_3N_2O_3$	Solid Foam, 52-54
262	$C_{32}H_{31}F_3N_2O_3$	Semi-Solid, 54-57
263	$C_{30}H_{29}F_6NO_5$	Oil/Solid, 55-65
264	$C_{30}H_{29}F_6NO_3$	Yellow Amorphous Solid
265	$C_{32}H_{28}F_6N_2O_3$	Solid Foam, 55-58
266	$C_{31}H_{32}F_6N_2O_5$	Liquid
267	$C_{30}H_{28}Cl_2N_2O_2$	Solid Foam 68-71
268	$C_{31}H_{28}F_4N_2O_3$	Solid Foam, 60-63
269	$C_{32}H_{29}F_5N_2O_4$	Solid Foam, 54-56
270	$C_{28}H_{26}F_6N_2O_2$	Solid

Cmpd No.	Empirical Formula	Physical State / Melting Point (°C)
271	C ₃₃ H ₃₀ F ₆ N ₂ O ₄	Solid
272	C ₃₃ H ₃₀ F ₆ N ₂ O ₄	Solid
273	C ₃₂ H ₃₅ F ₆ NO ₅	Oil
274	C ₃₂ H ₂₉ F ₆ N ₃ O ₅	Solid
275	C ₃₀ H ₂₇ F ₆ N ₃ O ₄	Solid
276	C ₃₃ H ₃₀ F ₆ N ₂ O ₄	Solid
277	C ₂₉ H ₂₇ ClF ₆ N ₂ O ₄	Liquid
278	C ₃₄ H ₃₀ ClF ₆ NO ₄	Solid Foam, 67-70
279	C ₃₁ H ₃₁ F ₆ NO ₆	Solid
280	C ₃₃ H ₃₅ F ₆ NO ₅	Solid, 63-68
281	C ₃₃ H ₂₇ F ₆ N ₃ O ₄	Solid
282	C ₃₃ H ₃₀ F ₆ N ₂ O ₄	Solid Foam, 53-56
283	C ₃₃ H ₂₇ F ₆ N ₃ O ₄	Solid
284	C ₄₂ H ₄₆ F ₃ N ₃ O ₄	Oil
285	C ₂₉ H ₂₈ F ₆ N ₂ O ₅	Solid, 83-92
286	C ₃₃ H ₂₈ ClF ₆ NO ₄	Syrup
287	C ₃₃ H ₃₃ F ₃ N ₂ O ₄	Solid Foam, 50-53
288	C ₃₁ H ₂₉ F ₃ N ₂ O ₃	Solid Foam, 49-53
289	C ₃₆ H ₃₂ ClF ₆ NO ₅	Solid Foam, 58-64
290	C ₃₃ H ₂₈ ClF ₆ NO ₄	Syrup
291	C ₃₁ H ₃₁ F ₆ NO ₄	Solid
292	C ₃₁ H ₃₁ F ₆ NO ₅	Solid, 68-75
293	C ₃₀ H ₂₇ Cl ₂ F ₆ NO ₄	Oil
294	C ₃₅ H ₃₄ F ₆ N ₂ O ₆	Glass
295	C ₃₄ H ₃₃ F ₃ N ₂ O ₅	Solid Foam, 52-56
296	C ₃₀ H ₂₇ F ₆ N ₃ O ₄	Liquid
297	C ₃₃ H ₂₈ ClF ₆ NO ₄	Solid, 45-47
298	C ₃₁ H ₃₀ F ₆ N ₂ O ₅	Semi Solid
299	C ₃₁ H ₃₁ F ₆ NO ₂	Solid
300	C ₃₁ H ₃₂ F ₆ N ₂ O ₅	
301	C ₃₀ H ₂₇ F ₉ N ₂ O ₄	Oil
302	C ₂₇ H ₂₅ F ₆ NO ₂	Solid
303	C ₃₀ H ₃₁ F ₆ NO ₂	Solid
304	C ₃₄ H ₃₁ F ₆ NO ₄	Solid
305	C ₃₈ H ₄₂ F ₆ N ₂ O ₄	Syrup
306	C ₃₅ H ₂₉ F ₁₂ N ₃ O ₄	Solid
307	C ₃₁ H ₂₆ ClF ₆ N ₃ O ₂	Solid
308	C ₃₀ H ₃₁ F ₆ NO ₅	Glass
309	C ₃₂ H ₂₈ F ₆ N ₂ O ₃	Solid
310	C ₃₁ H ₃₂ F ₆ N ₂ O ₅	Oil
311	C ₃₂ H ₃₄ F ₆ N ₂ O ₅	
312	C ₃₀ H ₂₉ F ₇ N ₂ O ₄	Oil
313	C ₃₁ H ₃₂ F ₆ N ₂ O ₃	Solid
314	C ₃₁ H ₂₈ ClF ₃ N ₂ O ₃	Solid, 61-85/Solid Foam, 65-95
315	C ₃₁ H ₃₁ F ₆ NO ₄	Solid
316	C ₃₄ H ₃₁ F ₆ NO ₂	Solid
317	C ₃₂ H ₃₂ F ₆ N ₂ O ₇	
318	C ₃₀ H ₃₀ F ₆ N ₂ O ₃	Solid
319	C ₃₂ H ₃₂ F ₆ N ₂ O ₃	Oil
320	C ₃₃ H ₃₄ F ₆ N ₂ O ₅	
321	C ₃₃ H ₃₀ F ₆ N ₂ O ₄	Solid
322	C ₃₂ H ₂₇ ClF ₆ N ₂ O ₄	Solid
323	C ₃₅ H ₃₀ Cl ₂ F ₆ N ₂ O ₅	
324	C ₃₄ H ₂₉ ClF ₅ N ₂ O ₅	
325	C ₃₀ H ₂₇ ClF ₃ N ₃ O ₃	Solid, 75-81
326	C ₃₂ H ₃₄ F ₆ N ₂ O ₅	
327	C ₃₂ H ₃₂ F ₆ N ₂ O ₅	Solid

Cmpd No.	Emperical Formula	Physical State / Melting Point (°C)
328	C ₂₉ H ₃₀ ClF ₃ N ₂ O ₃	Solid Foam, 54-57
329	C ₃₄ H ₃₆ F ₆ N ₂ O ₅	
330	C ₃₁ H ₃₁ F ₆ NO ₃	Solid
331	C ₂₇ H ₂₅ F ₆ NO ₄	Solid
335	C ₃₅ H ₃₇ F ₃ N ₂ O ₄	Solid, 108-112
336	C ₃₃ H ₃₆ F ₆ N ₂ O ₇	Solid, 137-142
337	C ₃₁ H ₃₂ F ₆ N ₂ O ₆	Solid, 173-176
338	C ₃₂ H ₃₁ F ₃ N ₂ O ₅	Solid, 98-104
339	C ₃₁ H ₂₈ BrF ₃ N ₂ O ₄	Solid, 103-107
340	C ₃₂ H ₃₁ F ₃ N ₂ O ₄	Solid, 85-90
341	C ₃₀ H ₂₉ F ₆ NO ₆	Solid, 164-185
342	C ₃₀ H ₂₉ F ₆ NO ₄	Yellow Solid
343	C ₃₂ H ₂₈ F ₆ N ₂ O ₄	Solid, 118-121
344	C ₃₁ H ₃₂ F ₆ N ₂ O ₆	Solid, 165-174
345	C ₃₀ H ₂₈ Cl ₂ N ₂ O ₃	Solid, 95-101
346	C ₃₁ H ₂₈ F ₄ N ₂ O ₄	Solid, 94-96
347	C ₃₂ H ₂₉ F ₅ N ₂ O ₅	Solid, 98-118
349	C ₃₃ H ₃₀ F ₆ N ₂ O ₅	Solid
350	C ₃₃ H ₃₀ F ₆ N ₂ O ₅	Solid
351	C ₃₂ H ₃₅ F ₆ NO ₆	Solid
353	C ₃₀ H ₂₇ F ₆ N ₃ O ₅	Solid, 136-140
354	C ₃₃ H ₃₀ F ₆ N ₂ O ₅	Solid
355	C ₂₉ H ₂₇ ClF ₆ N ₂ O ₅	Solid, 181-184
357	C ₃₁ H ₃₁ F ₆ NO ₇	Solid
358	C ₃₃ H ₃₅ F ₆ NO ₆	Solid, 210-212
359	C ₃₃ H ₂₇ F ₆ N ₃ O ₅	Solid
370	C ₃₁ H ₃₁ F ₆ NO ₆	Solid, 212-214
371	C ₃₀ H ₂₇ Cl ₂ F ₆ NO ₅	Oil
372	C ₃₅ H ₃₄ F ₆ N ₂ O ₇	Solid
375	C ₃₃ H ₂₈ ClF ₆ NO ₅	Solid Foam, 70-74
378	C ₃₀ H ₂₇ F ₉ N ₂ O ₅	Solid, 192-194
380	C ₃₀ H ₃₁ F ₆ NO ₃	Solid
381	C ₃₄ H ₃₁ F ₆ NO ₅	Solid
383	C ₃₁ H ₃₁ F ₆ NO ₃	Solid
384	C ₃₁ H ₂₆ ClF ₆ N ₃ O ₃	Solid
385	C ₃₀ H ₃₁ F ₆ NO ₆	Solid
387	C ₃₁ H ₃₂ F ₆ N ₂ O ₆	Oil
389	C ₃₀ H ₂₉ F ₇ N ₂ O ₅	Solid, 193-195
390	C ₃₁ H ₃₂ F ₆ N ₂ O ₄	Solid
391	C ₃₁ H ₂₈ ClF ₃ N ₂ O ₄	Solid, 62-66
392	C ₃₁ H ₃₁ F ₆ NO ₅	Solid
393	C ₃₄ H ₃₁ F ₆ NO ₃	Solid
395	C ₃₀ H ₃₀ F ₆ N ₂ O ₄	Solid
396	C ₃₂ H ₃₂ F ₆ N ₂ O ₄	Solid
399	C ₃₂ H ₂₇ ClF ₆ N ₂ O ₅	Solid, 200-203
402	C ₃₀ H ₂₇ ClF ₃ N ₃ O ₄	Solid, 84-88
404	C ₃₂ H ₃₂ F ₆ N ₂ O ₆	Solid
405	C ₂₉ H ₃₀ ClF ₃ N ₂ O ₄	Solid, 77-82
407	C ₃₁ H ₃₁ F ₆ NO ₄	Solid
409	C ₃₅ H ₂₉ F ₁₂ N ₃ O ₅	Solid
410	C ₃₀ H ₂₉ F ₆ N ₃ O ₆	Solid, 210-213
411	C ₃₂ H ₂₇ ClF ₆ N ₂ O ₄	Solid, 204-207
412	C ₃₁ H ₂₆ ClF ₆ N ₃ O ₅	Solid, 210-212
413	C ₃₂ H ₂₉ F ₆ N ₃ O ₅	Solid
416	C ₃₀ H ₂₉ F ₆ N ₅ O ₄	Solid
417	C ₃₀ H ₃₀ F ₆ N ₂ O ₅	Solid
418	C ₂₈ H ₂₅ F ₆ NO ₅	Solid

Cmpd No.	Emperical Formula	Physical State / Melting Point (°C)
419	$C_{31}H_{32}F_6N_2O_6$	Solid
421	$C_{32}H_{28}F_6N_2O_6$	Solid
422	$C_{31}H_{27}F_6N_3O_6$	Solid
423	$C_{31}H_{32}F_6N_2O_7$	Solid
424	$C_{30}H_{30}F_6N_2O_6$	Solid
425	$C_{31}H_{26}ClF_6N_3O_6$	Solid
426	$C_{30}H_{29}F_6N_5O_5$	Solid
430	$C_{32}H_{27}F_7N_2O_5$	Solid
431	$C_{32}H_{26}F_6N_2O_5$	Solid
432	$C_{31}H_{25}F_6N_3O_3$	Solid
433	$C_{31}H_{27}Cl_2F_3N_2O_3$	Solid Foam, 69-71
434	$C_{29}H_{27}F_2N_3O_2$	Solid Foam
435	$C_{32}H_{30}ClF_3N_2O_3$	Solid Foam, 60-64
436	$C_{32}H_{30}ClF_3N_2O_3$	Solid Foam, 59-62
437	$C_{31}H_{28}ClF_3N_2O_3$	Solid Foam, 59-62
438	$C_{31}H_{27}Cl_2F_3N_2O_3$	Solid Foam, 59-63
439	$C_{31}H_{28}ClF_3N_2O_3$	Solid Foam, 55-58
440	$C_{34}H_{34}F_3N_3O_4$	Solid Foam, 53-56
441	$C_{31}H_{27}Cl_2F_3N_2O_3$	Solid Foam, 60-64
442	$C_{31}H_{27}Cl_2F_3N_2O_3$	Solid Foam, 58-62
443	$C_{31}H_{27}Cl_2F_3N_2O_3$	Solid Foam, 55-59
444	$C_{29}H_{29}Cl_2F_3N_2O_3$	Solid Foam, 55-59
445	$C_{31}H_{27}ClF_4N_2O_3$	Solid Foam, 60-63
446	$C_{32}H_{26}F_4N_2O_6$	Oil/Solid Foam, 60-64
447	$C_{33}H_{30}F_6N_2O_5$	Solid
448	$C_{32}H_{29}F_6N_3O_2$	Solid
449	$C_{32}H_{34}F_6N_4O$	Solid
450	$C_{32}H_{34}F_6N_2O$	Solid
451	$C_{34}H_{32}F_6N_4O_2$	Solid
452	$C_{34}H_{31}F_6N_3O_3$	Solid
453	$C_{29}H_{29}Cl_2F_3N_2O_4$	Solid, 83-86
454	$C_{31}H_{27}Cl_2F_3N_2O_4$	Solid, 101-105
455	$C_{31}H_{27}Cl_2F_3N_2O_4$	Solid, 100-104
456	$C_{31}H_{27}Cl_2F_3N_2O_4$	Solid, 79-82
457	$C_{31}H_{27}Cl_2F_3N_2O_4$	Solid, 98-102
458	$C_{31}H_{28}ClF_3N_2O_4$	Solid, 85-89
459	$C_{31}H_{28}ClF_3N_2O_4$	Solid, 98-102
460	$C_{31}H_{27}Cl_2F_3N_2O_4$	Solid Foam, 76-80
461	$C_{32}H_{30}ClF_3N_2O_4$	Solid, 145-149
462	$C_{32}H_{30}ClF_3N_2O_4$	Solid, 89-92
463	$C_{31}H_{27}ClF_4N_2O_4$	Solid, 84-87
464	$C_{32}H_{26}F_4N_2O_7$	Solid Foam, 72-77
465	$C_{33}H_{30}F_6N_2O_6$	Solid
466	$C_{34}H_{32}F_6N_4O_3$	Solid
467	$C_{32}H_{32}F_6N_4O_2$	Solid
468	$C_{34}H_{31}F_6N_3O_4$	Solid
470	$C_{29}H_{28}ClF_6NO_5$	Solid Foam, 42-45
471	$C_{29}H_{28}ClF_6NO_5$	Solid Foam, 42-44
472	$C_{29}H_{28}ClF_6NO_5$	Solid Foam, 43-45
473	$C_{29}H_{28}ClF_6NO_4$	Syrup
474	$C_{28}H_{26}ClF_6NO_3$	Syrup
475	$C_{28}H_{26}ClF_6NO_4$	Syrup
476	$C_{30}H_{30}ClF_6NO_4$	Oil
477	$C_{28}H_{26}BrF_6NO_4$	Syrup
478	$C_{28}H_{26}ClF_6NO_5$	Solid, 54-58
479	$C_{28}H_{26}ClF_6NO_4$	Solid, 55-60
480	$C_{30}H_{30}ClF_6NO_5$	Semi-Solid

Cmpd No.	Emperical Formula	Physical State / Melting Point (°C)
481	C ₂₉ H ₂₈ ClF ₆ NO ₅	Solid, 57-61
482	C ₂₈ H ₂₆ BrF ₆ NO ₅	Solid, 60-65
483	C ₃₀ H ₃₁ F ₆ N ₂ O ₅ .C ₃ H ₇ O ₄ S	Solid, 120-128
484	C ₃₀ H ₃₁ F ₆ N ₂ O ₅ .C ₅ H ₉ O ₃	Solid, 74-80
485	C ₃₁ H ₂₇ ClF ₆ N ₃ O ₅ .C ₈ H ₁₅ O ₂	Solid, 190-194
486	C ₃₀ H ₃₁ F ₆ N ₂ O ₅ .C ₈ H ₁₅ O ₂	Solid, 53-65
487	C ₃₀ H ₃₁ F ₆ N ₂ O ₅ .C ₄ F ₉ O ₃ S	Solid, 85-94
488	C ₃₁ H ₂₇ ClF ₆ N ₃ O ₅ .C ₆ H ₇ O ₇	Solid, 132-142
489	C ₃₀ H ₃₀ F ₆ N ₃ O ₆ .C ₆ H ₇ O ₇	Solid, 113-123
490	C ₃₀ H ₃₁ F ₆ N ₂ O ₅ .C ₁₀ H ₁₅ O ₄ S	Solid, 123-131
491	C ₃₀ H ₃₁ F ₆ N ₂ O ₅ .C ₂ H ₅ O ₃ S	Solid, 188-192
492	C ₃₁ H ₂₇ ClF ₆ N ₃ O ₅ .C ₆ H ₇ O ₇	Solid, 135-144
493	C ₃₀ H ₃₁ F ₆ N ₂ O ₅ .C ₇ H ₁₁ O ₂	Solid, 78-88
494	C ₃₀ H ₃₁ F ₆ N ₂ O ₅ .C ₆ H ₇ O ₇	Solid, 114-119
495	C ₃₀ H ₃₁ F ₆ N ₂ O ₅ .C ₈ F ₁₇ O ₃ S	Solid, 74-84
496	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₈ F ₁₇ O ₃ S	Solid, 96-101
497	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .Cl	Solid, 140-143
498	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₁₀ H ₁₅ O ₄ S	Solid, 122-136
499	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₈ H ₁₅ O ₂	Solid, 55-65
500	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₃ H ₇ O ₄ S	Solid, 111-128
501	C ₂₉ H ₂₉ ClF ₆ N ₂ O ₅ .C ₆ H ₇ O ₇	Solid, 96-115
502	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₇ H ₁₁ O ₂	Solid, 80-85
503	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₅ H ₉ O ₃	Solid, 75-81
504	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₆ H ₇ O ₇	Solid, 195-197
505	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₁₀ H ₁₅ O ₄ S	Solid, 121-132
506	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₂ H ₅ O ₃ S	Solid, 168-184
507	C ₃₀ H ₃₁ F ₆ N ₂ O ₅ .C ₁₀ H ₁₅ O ₄ S	Solid, 118-119
508	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₄ F ₉ O ₃ S	Solid, 94-104
509	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₁₀ H ₁₇ O ₂	Solid, 43-58
510	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₂ H ₃ O ₃	Solid, 87-103
511	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₁₁ H ₁₅ O ₂	Solid, 100-125
512	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₅ H ₇ O ₄	Solid, 80-85
513	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₁₈ H ₃₃ O ₂	Liquid
514	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₇ H ₁₁ O ₄	Solid, 78-88
515	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₁₈ H ₂₉ O ₃ S	Solid, 85-97
516	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₆ H ₉ O ₄	Solid, 78-90
517	C ₃₂ H ₂₉ F ₆ N ₂ O ₅ .C ₈ H ₁₅ O ₂	Solid, 44-65
518	C ₃₁ H ₂₇ ClF ₆ N ₃ O ₅ .C ₂ H ₅ O ₃ S	Solid
519	C ₃₁ H ₂₇ ClF ₆ N ₃ O ₅ .C ₉ H ₉ ClF ₃ O ₂	Solid, 112-117
520	C ₃₁ H ₂₇ ClF ₆ N ₃ O ₅ .C ₈ F ₁₇ O ₃ S	Solid
521	C ₄₂ H ₄₆ F ₆ N ₃ O ₈ .Cl	Solid,
522	C ₃₂ H ₂₉ F ₆ N ₂ O ₄ .ClH	Solid, 78-82

Candidate insecticides were evaluated for activity against the tobacco budworm (*Heliothis virescens* [Fabricius]) in a surface-treated diet test.

- 5 In this test one mL of molten (65-70°C) wheat germ-based artificial diet was pipetted into each well of a four by six (24 well) multi-well plate (ID# 430345-15.5 mm dia. x 17.6 mm deep; Corning Costar Corp., One Alewife Center, Cambridge, MA 02140). The diet was allowed to cool to ambient temperature before treatment with candidate insecticide.

For a determination of insecticidal activity, solutions of the candidate insecticides were prepared for testing using a Packard 204DT Multiprobe[®] Robotic System (Packard Instrument Company, 800 Research Parkway, Meriden, CT 06450), in which the robot first diluted a standard 50 millimolar DMSO solution of candidate insecticide with a 1:1 water/acetone solution (V/V) in a ratio of 1:7 stock solution to water/acetone. The robot subsequently pipetted 40 microliters of the so-prepared solution onto the surface of the diet in each of three wells in the 24 multi-well plate. The process was repeated with solutions of seven other candidate insecticides. Once treated, the contents of the multi-well plate were allowed to dry, leaving 0.25 millimoles of candidate insecticide on the surface of the diet, or a concentration of 0.25 millimolar. Appropriate untreated controls containing only DMSO on the diet surface were also included in this test.

For evaluations of the insecticidal activity of a candidate insecticide at varying rates of application, the test was established as described above using sub-multiples of the standard 50 millimolar DMSO solution of candidate insecticide. For example, the standard 50 millimolar solution was diluted by the robot with DMSO to give 5, 0.5, 0.05, 0.005, 0.0005 millimolar, or more dilute solutions of the candidate insecticide. In these evaluations there were six replicates of each rate of application placed on the surface of the diet in the 24 multi-well plate, for a total of four rates of application of candidate insecticide in each plate.

In each well of the test plate was placed one second instar tobacco budworm larva, each weighing approximately five milligrams. After the larvae were placed in each well, the plate was sealed with clear polyfilm adhesive tape. The tape over each well was perforated to ensure an adequate air supply. The plates were then held in a growth chamber at 25 °C and 60% relative humidity for five days (light 14 hours/day).

After the five-day exposure period insecticidal activity for each rate of application of candidate insecticide was assessed as percent inhibition of insect weight relative to the weight of insects from untreated controls, and percent mortality when compared to the total number of insects infested.

Insecticidal activity data at selected rates of application from this test are provided in Table 3. The test compounds of formula I are identified by numbers that correspond to those in Table 1.

Table 3
Insecticidal Activity of Test Compounds Applied to the
Surface of the Diet of Tobacco Budworm

5

[illegible]

[illegible]

[illegible]

Inhibition										
Cmpd. No	512	513	514	515	516	517	518	519	520	521
Percent Mortality	100	100	100	100	100	100	100	100	100	50
Percent Growth Inhibition	100	100	100	100	100	100	100	100	100	100
Cmpd. No	522									
Percent Mortality	100									
Percent Growth Inhibition	100									

These tests were conducted with 0.25 millimoles of candidate insecticide on the surface of the diet

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As set forth in the foregoing Table 3, most of the compounds therein provided 100% mortality and 100% growth inhibition of tobacco budworm.

While this invention has been described with an emphasis upon preferred embodiments, it will be understood by those of ordinary skill in the art that variations of the preferred embodiments may be used and that it is intended that the invention may be practiced otherwise than as specifically described herein. Accordingly, this invention includes all modifications encompassed within the spirit and scope of the invention as defined by the following claims.

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